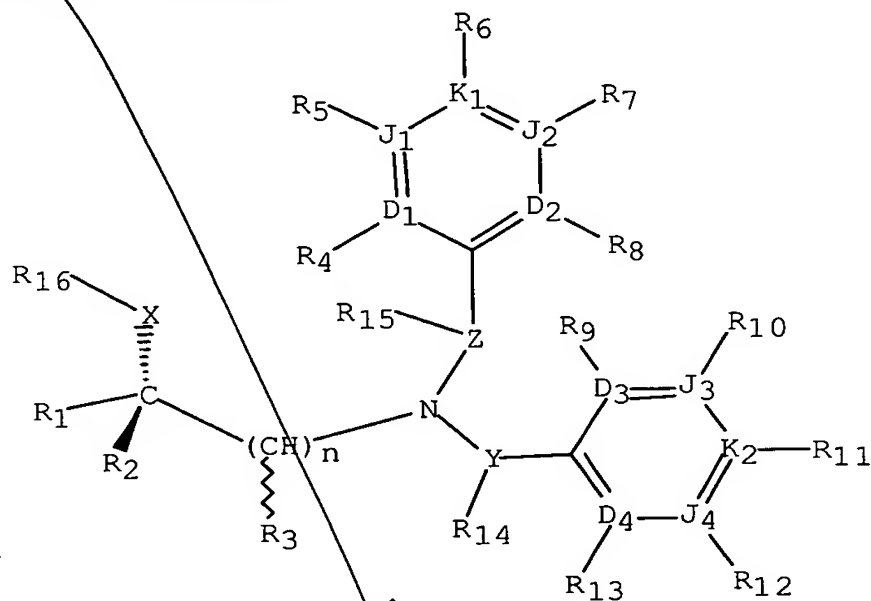


What we claim is:

1. A compound having the formula:



5

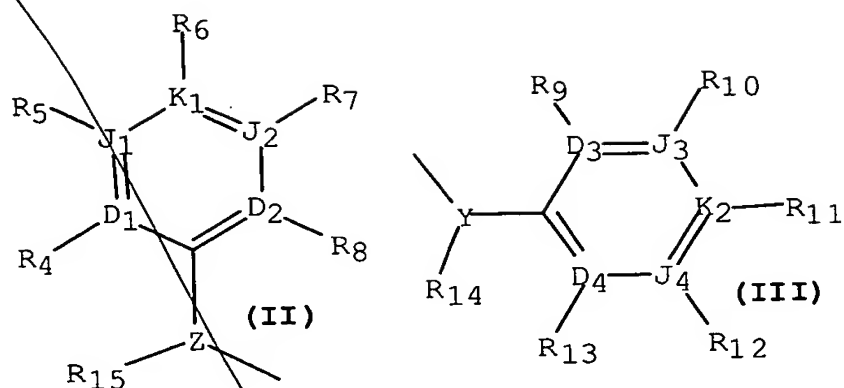
and a pharmaceutically-acceptable salt thereof, wherein;

n is an integer selected from 1 through 4;

X is oxy;

R_1 is selected from the group consisting of haloalkyl, haloalkenyl,

10 haloalkoxymethyl, and haloalkenyloxymethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(A)Q$ wherein A is Formula (II) and Q is Formula (III);



R₁₆ is selected from the group consisting of hydrido, alkyl, acyl, aroyl, heteroaroyl, trialkylsilyl, and a spacer selected from the group consisting of a covalent single bond and a linear spacer moiety having a chain length of 1 to 4 atoms linked to the point of bonding of any aromatic substituent selected from the group consisting of R₄, R₈, R₉, and R₁₃ to form a heterocyclyl ring having from 5 through 10 contiguous members;

D₁, D₂, J₁, J₂ and K₁ are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that no more than one of D₁, D₂, J₁, J₂ and K₁ is a covalent bond, no more than one of D₁, D₂, J₁, J₂ and K₁ is O, no more than one of D₁, D₂, J₁, J₂ and K₁ is S, one of D₁, D₂, J₁, J₂ and K₁ must be a covalent bond when two of D₁, D₂, J₁, J₂ and K₁ are O and S, and no more than four of D₁, D₂, J₁, J₂ and K₁ are N;

D₃, D₄, J₃, J₄ and K₂ are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that no more than one is a covalent bond, no more than one of D₃, D₄, J₃, J₄ and K₂ is O, no more than one of D₃, D₄, J₃, J₄ and K₂ is S, no more than two of D₃, D₄, J₃, J₄ and K₂ are O and S; one of D₃, D₄, J₃, J₄ and K₂ must be a covalent

bond when two of D_3 , D_4 , J_3 , J_4 and K_2 are O and S, and no more than four of D_3 , D_4 , J_3 , J_4 and K_2 are N;

R_2 is selected from the group consisting of hydrido, aryl, aralkyl, alkyl, alkenyl, alkenyloxyalkyl, haloalkyl, haloalkenyl, halocycloalkyl, haloalkoxy, haloalkoxyalkyl, haloalkenyloxyalkyl, halocycloalkoxy, halocycloalkoxyalkyl, perhaloaryl, perhaloaralkyl, perhaloaryloxyalkyl, heteroaryl, dicyanoalkyl, and carboalkoxycyanoalkyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

R_3 is selected from the group consisting of hydrido, hydroxy, cyano, aryl, aralkyl, acyl, alkoxy, alkyl, alkenyl, alkoxyalkyl, heteroaryl, alkenyloxyalkyl, haloalkyl, haloalkenyl, haloalkoxy, haloalkoxyalkyl, haloalkenyloxyalkyl, monocyanalkyl, dicyanoalkyl, carboxamide, and carboxamidoalkyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from a group consisting of a covalent single bond, $(C(R_{14})_2)_q$ wherein q is an integer selected from 1 and 2 and $(CH(R_{14}))_g-W-(CH(R_{14}))_p$ wherein g and p are integers independently selected from 0 and 1;

R_{14} is selected from the group consisting of hydrido, hydroxy, cyano, hydroxyalkyl, acyl, alkoxy, alkyl, alkenyl, alkynyl, alkoxyalkyl, haloalkyl, haloalkenyl, haloalkoxy, haloalkoxyalkyl, haloalkenyloxyalkyl, monocarboalkoxyalkyl, monocyanoalkyl, dicyanoalkyl, carboalkoxycyanoalkyl, carboalkoxy, carboxamide, carboxamidoalkyl;

Z is selected from the group consisting of covalent single bond,
 $(C(R_{15})_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH(R_{15}))_j-W-$
 $(CH(R_{15}))_k$ wherein j and k are integers independently selected from 0 and 1;

5 W is selected from the group consisting of O, C(O), C(S),
 C(O)N(R₁₄), C(S)N(R₁₄), (R₁₄)NC(O), (R₁₄)NC(S), S, S(O), S(O)₂,
 S(O)₂N(R₁₄), (R₁₄)NS(O)₂, and N(R₁₄) with the proviso that R₁₄ is other
 than cyano;

R₁₅ is selected from the group consisting of hydrido, cyano,
 10 hydroxyalkyl, acyl, alkoxy, alkyl, alkenyl, alkynyl, alkoxyalkyl, haloalkyl,
 haloalkenyl, haloalkoxy, haloalkoxyalkyl, haloalkenyloxyalkyl,
 monocarboalkoxyalkyl, monocyanoalkyl, dicyanoalkyl,
 carboalkoxycyanoalkyl, carboalkoxy, carboxamide, and carboxamidoalkyl;

R₄, R₅, R₆, R₇, R₈, R₉, R₁₀, R₁₁, R₁₂, and R₁₃ are independently
 15 selected from the group consisting of hydrido, carboxy, heteroaralkylthio,
 heteroaralkoxy, cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy,
 heterocyclyloxy, aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocyclyl,
 perhaloaralkyl, aralkylsulfonyl, aralkylsulfonylalkyl, aralkylsulfinyl,
 aralkylsulfinylalkyl, halocycloalkyl, halocycloalkenyl, cycloalkylsulfinyl,
 20 cycloalkylsulfinylalkyl, cycloalkylsulfonyl, cycloalkylsulfonylalkyl,
 heteroaryl amino, N-heteroaryl amino-N-alkyl amino,
 heteroaryl aminoalkyl, haloalkylthio, alkanoyloxy, alkoxy, alkoxyalkyl,
 haloalkoxylalkyl, heteroaralkoxy, cycloalkoxy, cycloalkenyloxy,
 cycloalkoxyalkyl, cycloalkylalkoxy, cycloalkenyloxyalkyl,
 25 cycloalkylenedioxy, halocycloalkoxy, halocycloalkoxyalkyl,
 halocycloalkenyloxy, halocycloalkenyloxyalkyl, hydroxy, amino, thio, nitro,
 lower alkyl amino, alkylthio, alkylthioalkyl, aryl amino, aralkyl amino,

arylthio, arylthioalkyl, heteroaralkoxyalkyl, alkylsulfinyl, alkylsulfinylalkyl,
 arylsulfinylalkyl, arylsulfonylalkyl, heteroarylsulfinylalkyl,
 heteroarylsulfonylalkyl, alkylsulfonyl, alkylsulfonylalkyl,
 haloalkylsulfinylalkyl, haloalkylsulfonylalkyl, alkylsulfonamido,
 5 alkylaminosulfonyl, amidosulfonyl, monoalkyl amidosulfonyl, dialkyl
 amidosulfonyl, monoarylamidosulfonyl, arylsulfonamido,
 diarylamidosulfonyl, monoalkyl monoaryl amidosulfonyl, arylsulfinyl,
 arylsulfonyl, heteroarylthio, heteroarylsulfinyl, heteroarylsulfonyl,
 heterocyclisulfonyl, heterocyclylthio, alkanoyl, alkenoyl, aroyl, heteroaroyl,
 10 aralkanoyl, heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl, alkynyl,
 alkenyloxy, alkenyloxyalkyl, alkylenedioxy, haloalkylenedioxy, cycloalkyl,
 cycloalkylalkanoyl, cycloalkenyl, lower cycloalkylalkyl, lower
 cycloalkenylalkyl, halo, haloalkyl, haloalkenyl, haloalkoxy,
 hydroxyhaloalkyl, hydroxyaralkyl, hydroxyalkyl, hydroxyheteroaralkyl,
 15 haloalkoxyalkyl, aryl, heteroaralkynyl, aryloxy, aralkoxy, aryloxyalkyl,
 saturated heterocyclyl, partially saturated heterocyclyl, heteroaryl,
 heteroaryloxy, heteroaryloxyalkyl, arylalkenyl, heteroarylalkenyl,
 carboxyalkyl, carboalkoxy, alkoxycarboxamido, alkylamidocarbonylamido,
 arylamidocarbonylamido, carboalkoxyalkyl, carboalkoxyalkenyl,
 20 carboaralkoxy, carboxamido, carboxamidoalkyl, cyano, carbohaloalkoxy,
 phosphono, phosphonoalkyl, diaralkoxyphosphono, and
 diaralkoxyphosphonoalkyl with the proviso that and R_4 , R_5 , R_6 , R_7 , R_8 , R_9 ,
 R_{10} , R_{11} , R_{12} , and R_{13} are each independently selected to maintain the
 tetravalent nature of carbon, trivalent nature of nitrogen, the divalent nature
 25 of sulfur, and the divalent nature of oxygen;

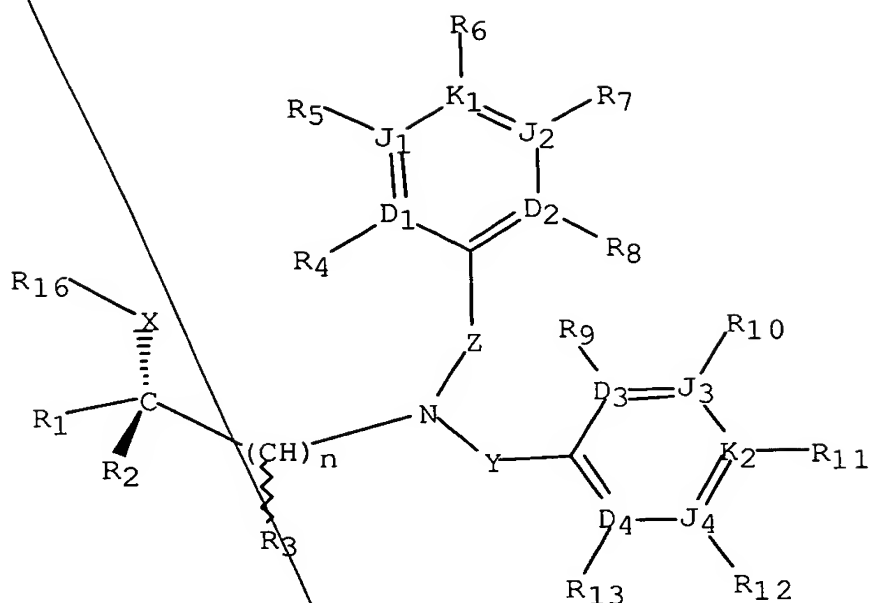
R_4 and R_5 , R_5 and R_6 , R_6 and R_7 , R_7 and R_8 , R_9 and R_{10} , R_{10} and

R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} are independently selected to form spacer
 pairs wherein a spacer pair is taken together to form a linear moiety having

from 3 through 6 contiguous atoms connecting the points of bonding of said spacer pair members to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 contiguous members, a partially saturated heterocyclyl ring having 5 through 8 contiguous members, a heteroaryl ring having 5 through 6 contiguous members, and an aryl with the provisos that no more than one of the group consisting of spacer pairs R₄ and R₅, R₅ and R₆, R₆ and R₇, and R₇ and R₈, is used at the same time and that no more than one of the group consisting of spacer pairs R₉ and R₁₀, R₁₀ and R₁₁, R₁₁ and R₁₂, and R₁₂ and R₁₃ is used at the same time;

10 R₄ and R₉, R₄ and R₁₃, R₈ and R₉, and R₈ and R₁₃ are independently selected to form a spacer pair wherein said spacer pair is taken together to form a linear moiety wherein said linear moiety forms a ring selected from the group consisting of a partially saturated heterocyclyl ring having from 5 through 8 contiguous members and a heteroaryl ring having from 5 through 6 contiguous members with the proviso that no more than one of the group consisting of spacer pairs R₄ and R₉, R₄ and R₁₃, R₈ and R₉, and R₈ and R₁₃ is used at the same time.

2. The compound as recited in Claim 1 having the formula of:



or a pharmaceutically acceptable salt thereof, wherein;

n is an integer selected from 1 through 3;

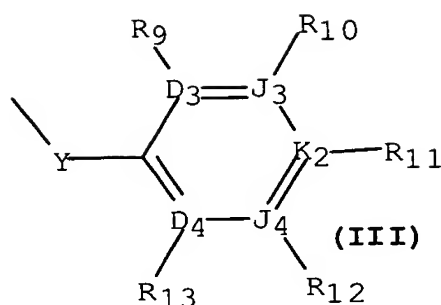
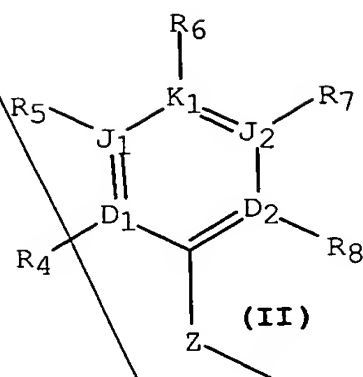
X is oxy;

R_{16} is selected from the group consisting of hydrido and a spacer selected from the group consisting of a covalent single bond and a linear spacer moiety having a chain length of 1 to 4 atoms linked to the point of bonding of any aromatic substituent selected from the group consisting of R_4 ,

R_8 , R_9 , and R_{13} to form a heterocycl ring having from 5 through 10 contiguous members;

R_1 is selected from the group consisting of haloalkyl and haloalkoxymethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CH_2)_n-N(A)Q$ wherein A

is Formula (II) and Q is Formula (III);



D_1, D_2, J_1, J_2 and K_1 are independently selected from the group consisting of C, N, O, S and a covalent bond with the provisos that no more than one of D_1, D_2, J_1, J_2 and K_1 is a covalent bond, no more than one of D_1, D_2, J_1, J_2 and K_1 is O, no more than one of D_1, D_2, J_1, J_2 and K_1 is S, one of D_1, D_2, J_1, J_2 and K_1 must be a covalent bond when two of D_1, D_2, J_1, J_2 and K_1 are O and S, and no more than four of D_1, D_2, J_1, J_2 and K_1 are N;

D_3, D_4, J_3, J_4 and K_2 are independently selected from the group consisting of C, N, O, S and a covalent bond with the provisos that no more than one of D_3, D_4, J_3, J_4 and K_2 is a covalent bond, no more than one of D_3, D_4, J_3, J_4 and K_2 is O, no more than one of D_3, D_4, J_3, J_4 and K_2 is S, one of D_3, D_4, J_3, J_4 and K_2 must be a covalent bond when two of D_3, D_4, J_3, J_4 and K_2 are O and S, and no more than four of D_3, D_4, J_3, J_4 and K_2 are N;

R_2 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, haloalkoxy, haloalkoxyalkyl, perhaloaryl, perhaloaralkyl,

perhaloaryloxyalkyl, and heteroaryl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

R_3 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, and haloalkoxyalkyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of a covalent single bond, $(CH_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH_2)_j-O-(CH_2)_k$ wherein j and k are integers independently selected from 0 and 1;

Z is selected from the group consisting of covalent single bond, $(CH_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH_2)_j-O-(CH_2)_k$ wherein j and k are integers independently selected from 0 and 1;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido, halo, haloalkyl, and alkyl ;

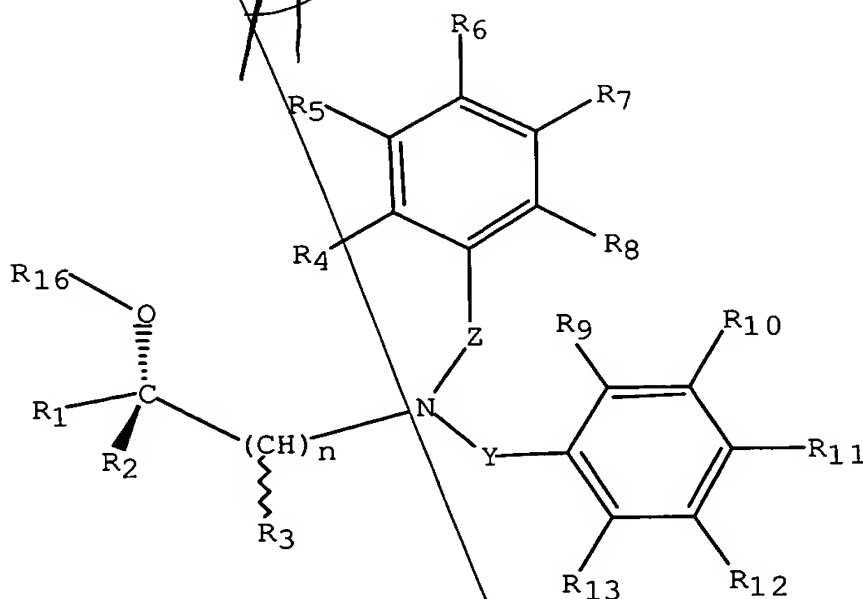
R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, carboxy, heteroaralkylthio, heteroaralkoxy, cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy, heterocyclyloxy, aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocyclyl, perhaloaralkyl, aralkylsulfonyl, aralkylsulfonylalkyl, aralkylsulfinyl, aralkylsulfinylalkyl, halocycloalkyl, halocycloalkenyl, cycloalkylsulfinyl, cycloalkylsulfinylalkyl, cycloalkylsulfonyl, cycloalkylsulfonylalkyl, heteroaryl amino, N-heteroaryl amino-N-alkyl amino, heteroaryl aminoalkyl, haloalkylthio, alkanoyloxy, alkoxy, alkoxyalkyl, haloalkoxylalkyl, heteroaralkoxy, cycloalkoxy, cycloalkenyloxy, cycloalkoxyalkyl, cycloalkylalkoxy, cycloalkenyloxyalkyl, cycloalkylenedioxy, halocycloalkoxy,

halocycloalkoxyalkyl, halocycloalkenyloxy, halocycloalkenyloxyalkyl, hydroxy, amino, thio, nitro, lower alkylamino, alkylthio, alkylthioalkyl, arylamino, aralkylamino, arylthio, arylthioalkyl, heteroaralkoxyalkyl, alkylsulfinyl, alkylsulfinylalkyl, arylsulfinylalkyl, arylsulfonylalkyl, heteroarylsulfinylalkyl, heteroarylsulfonylalkyl, alkylsulfonyl, alkylsulfonylalkyl, haloalkylsulfinylalkyl, haloalkylsulfonylalkyl, alkylsulfonamido, alkylaminosulfonyl, amidosulfonyl, monoalkyl amidosulfonyl, dialkyl amidosulfonyl, monoarylamidosulfonyl, arylsulfonamido, diarylamidosulfonyl, monoalkyl monoaryl amidosulfonyl, arylsulfinyl, arylsulfonyl, heteroarylthio, heteroarylsulfinyl, heteroarylsulfonyl, heterocyclylsulfonyl, heterocyclylthio, alkanoyl, alkenoyl, aroyl, heteroaroyl, aralkanoyl, heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl, alkynyl, alkenyloxy, alkenyloxyalkyl, alkylenedioxy, haloalkylenedioxy, cycloalkyl, cycloalkylalkanoyl, cycloalkenyl, lower cycloalkylalkyl, lower cycloalkenylalkyl, halo, haloalkyl, haloalkenyl, haloalkoxy, hydroxyhaloalkyl, hydroxyaralkyl, hydroxyalkyl, hydroxyheteroaralkyl, haloalkoxyalkyl, aryl, heteroaralkynyl, aryloxy, aralkoxy, aryloxyalkyl, saturated heterocyclyl, partially saturated heterocyclyl, heteroaryl, heteroaryloxy, heteroaryloxyalkyl, heteroaralkyl, arylalkenyl, heteroarylalkenyl, carboxyalkyl, carboalkoxy, alkoxycarboxamido, alkylamidocarbonylamido, arylamidocarbonylamido, carboalkoxyalkyl, carboalkoxyalkenyl, carboaralkoxy, carboxamido, carboxamidoalkyl, cyano, carbohaloalkoxy, phosphono, phosphonoalkyl, diaralkoxyphosphono, and diaralkoxyphosphonoalkyl;

R_4 and R_5 , R_5 and R_6 , R_6 and R_7 , R_7 and R_8 , R_9 and R_{10} , R_{10} and R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} are independently selected to form spacer pairs wherein a spacer pair is taken together to form a linear moiety having from 3 through 6 contiguous atoms connecting the points of bonding of said spacer pair members to form a ring selected from the group consisting of a

cycloalkenyl ring having 5 through 8 contiguous members, a partially saturated heterocyclyl ring having 5 through 8 contiguous members, a heteroaryl ring having 5 through 6 contiguous members, and an aryl with the provisos that no more than one of the group consisting of spacer pairs R_4 and R_5 , R_5 and R_6 , R_6 and R_7 , and R_7 and R_8 , is used at the same time and that no more than one of the group consisting of spacer pairs R_9 and R_{10} , R_{10} and R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} is used at the same time.

3. The compound as recited in Claim 2 having the formula of:



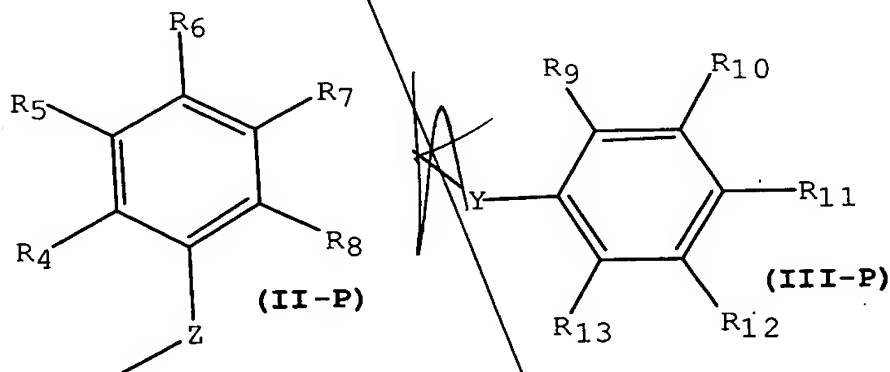
or a pharmaceutically acceptable salt thereof, wherein;

n is an integer selected from 1 through 3,

R_{16} is selected from the group consisting of hydrido and a spacer selected from the group consisting of a covalent single bond and a linear

spacer moiety having a chain length of 1 to 4 atoms linked to the point of bonding of any aromatic substituent selected from the group consisting of R_4 , R_8 , R_9 , and R_{13} to form a heterocyclyl ring having from 5 through 10 contiguous members;

- 5 R_1 is selected from the group consisting of haloalkyl and haloalkoxymethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



10

R_2 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, haloalkoxy, haloalkoxyalkyl, perhaloaryl, perhaloaralkyl, perhaloaryloxyalkyl, and heteroaryl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

- 15 R_3 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, and haloalkoxyalkyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking

than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of a covalent single bond, $(CH_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH_2)_j-O-(CH_2)_k$ wherein j and k are integers independently selected from 0 and 1;

Z is selected from the group consisting of covalent single bond, $(CH_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH_2)_j-O-(CH_2)_k$ wherein j and k are integers independently selected from 0 and 1;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido, halo, haloalkyl, and alkyl ;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, carboxy, heteroaralkylthio, heteroaralkoxy, cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy, heterocycloxy, aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocyclyl, perhaloaralkyl, aralkylsulfonyl, aralkylsulfonylalkyl, aralkylsulfinyl, aralkylsulfinylalkyl, halocycloalkyl, halocycloalkenyl, cycloalkylsulfinyl, cycloalkylsulfinylalkyl, cycloalkylsulfonyl, cycloalkylsulfonylalkyl, heteroaryl amino, N-heteroaryl amino-N-alkyl amino, heteroaryl aminoalkyl, haloalkylthio, alkanoyloxy, alkoxy, alkoxyalkyl, haloalkoxylalkyl, heteroaralkoxy, cycloalkoxy, cycloalkenyloxy, cycloalkoxyalkyl, cycloalkylalkoxy, cycloalkenyloxyalkyl, cycloalkylenedioxy, halocycloalkoxy, halocycloalkoxyalkyl, halocycloalkenyloxy, halocycloalkenyloxyalkyl, hydroxy, amino, thio, nitro, lower alkyl amino, alkylthio, alkylthioalkyl, aryl amino, aralkyl amino, arylthio, arylthioalkyl, heteroaralkoxyalkyl, alkylsulfinyl, alkylsulfinylalkyl, arylsulfinylalkyl, arylsulfonylalkyl, heteroaryl sulfinylalkyl, heteroaryl sulfonylalkyl, alkylsulfonyl, alkylsulfonylalkyl, haloalkylsulfinylalkyl, haloalkylsulfonylalkyl,

alkylsulfonamido, alkylaminosulfonyl, amidosulfonyl, monoalkyl
 amidosulfonyl, dialkyl amidosulfonyl, monoarylamidosulfonyl,
 arylsulfonamido, diarylamidosulfonyl, monoalkyl monoaryl amidosulfonyl,
 arylsulfinyl, arylsulfonyl, heteroarylthio, heteroarylsulfinyl,
 5 heteroarylsulfonyl, heterocyclisulfonyl, heterocyclylthio, alkanoyl, alkenoyl,
 aroyl, heteroaroyl, aralkanoyl, heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl,
 alkynyl, alkenyloxy, alkenyloxyalkyl, alkylenedioxy, haloalkylenedioxy,
 cycloalkyl, cycloalkylalkanoyl, cycloalkenyl, lower cycloalkylalkyl, lower
 cycloalkenylalkyl, halo, haloalkyl, haloalkenyl, haloalkoxy,
 10 hydroxyhaloalkyl, hydroxyaralkyl, hydroxyalkyl, hydroxyheteroaralkyl,
 haloalkoxyalkyl, aryl, heteroaralkynyl, aryloxy, aralkoxy, aryloxyalkyl,
 saturated heterocyclyl, partially saturated heterocyclyl, heteroaryl,
 heteroaryloxy, heteroaryloxyalkyl, heteroaralkyl, arylalkenyl,
 heteroarylalkenyl, carboxyalkyl, carboalkoxy, alkoxycarboxamido,
 15 alkylamidocarbonylamido, arylamidocarbonylamido, carboalkoxyalkyl,
 carboalkoxyalkenyl, carboalkoxy, carboxamido, carboxamidoalkyl, cyano,
 carbohaloalkoxy, phosphono, phosphonoalkyl, diaralkoxyphosphono, and
 diaralkoxyphosphonoalkyl;

R_4 and R_5 , R_5 and R_6 , R_6 and R_7 , R_7 and R_8 , R_9 and R_{10} , R_{10} and

20 R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} are independently selected to form spacer
 pairs wherein a spacer pair is taken together to form a linear moiety having
 from 3 through 6 contiguous atoms connecting the points of bonding of said
 spacer pair members to form a ring selected from the group consisting of a
 cycloalkenyl ring having 5 through 8 contiguous members, a partially
 25 saturated heterocyclyl ring having 5 through 8 contiguous members, a
 heteroaryl ring having 5 through 6 contiguous members, and an aryl with the
 provisos that no more than one of the group consisting of spacer pairs R_4 and
 R_5 , R_5 and R_6 , R_6 and R_7 , and R_7 and R_8 , is used at the same time and that

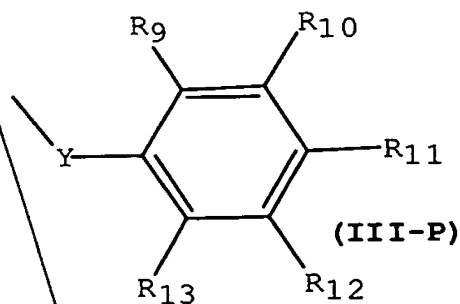
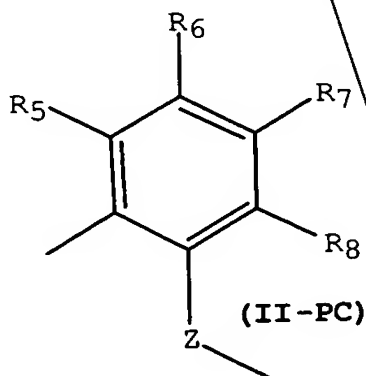
no more than one of the group consisting of spacer pairs R_9 and R_{10} , R_{10} and R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} is used at the same time.

4. The compound as recited in Claim 3 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

R_{16} is taken together with R_4 , R_8 , R_9 , or R_{13} to form a spacer selected from the group consisting of a covalent single bond, CH_2 , $CH(CH_3)$, CF_2 , $C(O)$, $C(S)$, and SO_2 ;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Apc)Qp$ wherein Apc is Formula (II-PC) and Qp is Formula (III-P);



;

R_2 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, vinyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Apc)Qp$;

R_3 is selected from the group consisting of hydrido, methyl, ethyl, vinyl, phenyl, 4-trifluoromethylphenyl, trifluoromethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the provisos that $(CHR_3)_n-N(Apc)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of covalent single bond, oxy, methyleneoxy, methylene, and ethylene;

Z is selected from the group consisting of covalent single bond, oxy, methyleneoxy, methylene, and ethylene;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 and R_{10} are independently selected from the group consisting of 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy, 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy,

- 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy,
 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy,
 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy,
 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl,
 5 cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl,
 cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy,
 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl,
 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy,
 3,4-difluorophenoxy, 2,3-difluorobenzoyloxy, 2,4-difluorobenzoyloxy,
 10 3,4-difluorobenzoyloxy, 2,5-difluorobenzoyloxy, 3,5-difluorophenoxy,
 3,4-difluorophenyl, 3,5-difluorobenzoyloxy, 4-difluoromethoxybenzoyloxy,
 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy,
 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy,
 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzoyloxy,
 15 3,5-dimethylbenzoyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl,
 1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy,
 4-ethylbenzoyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,
 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl,
 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzoyloxy,
 20 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzoyloxy,
 3-fluoro-5-trifluoromethylbenzoyloxy, 4-fluoro-2-trifluoromethylbenzoyloxy,
 4-fluoro-3-trifluoromethylbenzoyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzoyloxy,
 25 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
 3-iodobenzoyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzoyloxy,
 30 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,

- 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl,
 5-isothiazolyl, 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
 3-methoxycarbonylprop-2-enyloxy, 4-methoxyphenyl,
 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy,
 5 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl,
 pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 10 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy,
 phenylamino, 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy,
 propoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, *sec*-butyl,
 4-*sec*-butylphenoxy, *tert*-butoxy, 3-*tert*-butylphenoxy, 4-*tert*-butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 15 2-(5,6,7,8-tetrahydronaphthyl)oxy, thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy,
 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl,
 20 3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy,
 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,
 25 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, and
 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of chloro, fluoro, hydrido, difluoromethoxy, trifluoromethyl, trifluoromethoxy, pentafluoroethyl, and 1,1,2,2-tetrafluoroethoxy;

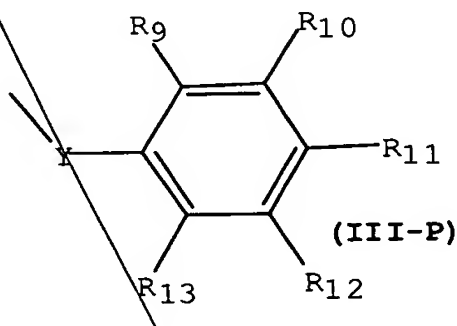
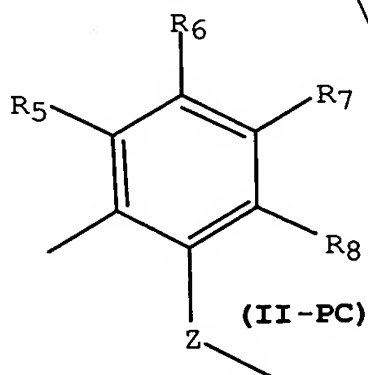
R_7 and R_{12} are independently selected from the group consisting of hydrido, fluoro, and trifluoromethyl.

5 The compound as recited in Claim 4 or a pharmaceutically acceptable salt thereof, wherein:

10 R_{16} is taken together with R_4 , R_8 , R_9 , or R_{13} to form a covalent single bond.

n is the integer 1;

R_1 is selected from the group consisting of trifluoromethyl and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Apc)Qp$ wherein
15 Apc is Formula (II-PC) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido and phenyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Apc)Qp$;

R_3 is selected from the group consisting of hydrido, methyl, trifluoromethyl, and difluoromethyl with the provisos that $(CHR_3)_n-N(Apc)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is methylene;
Z is covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy, 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of cyclopentyl, 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl, isobutyl, isopropoxy, pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of fluoro and hydrido;

R_7 and R_{12} are independently selected from the group consisting of hydrido and fluoro.

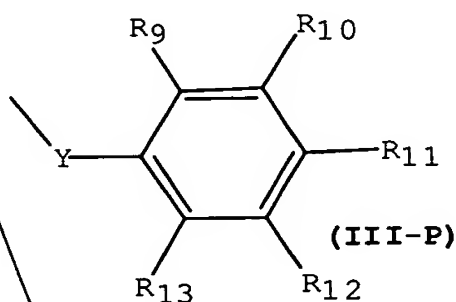
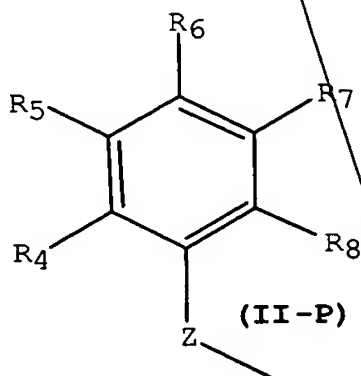
5

6. The compound as recited in Claim 3 or a pharmaceutically acceptable salt thereof, wherein:

10

n is an integer selected from 1 and 2;

R_1 is selected from the group consisting of haloalkyl and haloalkoxymethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



15

R_{16} is hydrido;

R_2 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, haloalkoxy, haloalkoxyalkyl, perhaloaryl, perhaloaralkyl,

perhaloaryloxyalkyl, and heteroaryl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, and haloalkoxyalkyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of a covalent single bond, oxy and C1-C2 alkylenes;

Z is a covalent single bond;

R_4, R_8, R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

$R_5, R_6, R_7, R_{10}, R_{11}$, and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, heterocyclyloxy, aralkylaryl, heteroaryloxyalkyl, heteroarylthio, and heteroarylsulfonyl.

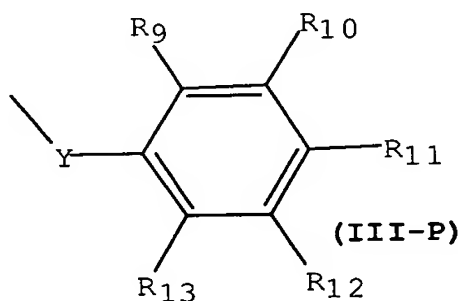
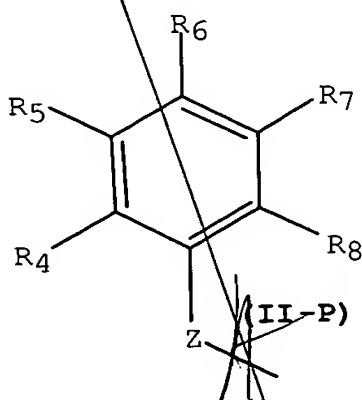
7. The compound as recited in Claim 6 or a pharmaceutically acceptable salt thereof wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and

5 $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, methyl, ethyl, propyl, butyl, vinyl, phenyl, 4-trifluoromethylphenyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, and

10 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, ethyl, vinyl, trifluoromethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and

15 pentafluoroethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is a covalent single bond;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

- 5 R_5 and R_{10} are independently selected from the group consisting of 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy, 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy,
- 10 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy,
- 15 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl, cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl, cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl,
- 20 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorobenzyloxy, 3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 3,5-difluorobenzyloxy, 4-difluoromethoxybenzyloxy, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy,
- 25 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl, 1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy, 4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,

- 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl,
 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy,
 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy,
 5 3-fluoro-5-trifluoromethylbenzyloxy, 4-fluoro-2-trifluoromethylbenzyloxy,
 4-fluoro-3-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy,
 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
 10 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
 3-iodobenzyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzyloxy,
 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl,
 15 5-isothiazolyl, 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
 3-methoxycarbonylprop-2-enyloxy, 4-methoxyphenyl,
 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy,
 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 20 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl,
 pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy,
 phenylamino, 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy,
 25 propoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, *sec*-butyl,
 4-*sec*-butylphenoxy, *tert*-butoxy, 3-*tert*-butylphenoxy, 4-*tert*-butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy,
 30 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,

3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl,
 3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy,
 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 5 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,
 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
 10 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, and
 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy,
 15 trifluoromethyl, and trifluoromethoxy;

R_7 and R_{12} are independently selected from the group consisting of
 hydrido, fluoro, and trifluoromethyl.

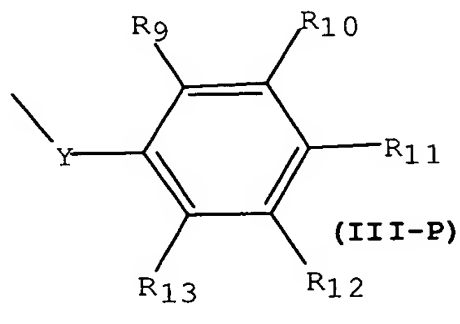
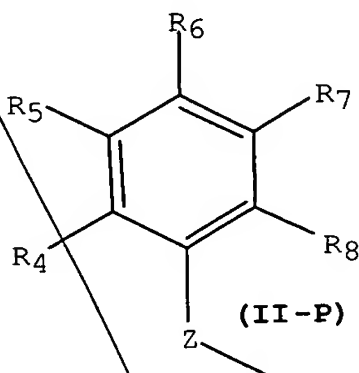
8. The compound as recited in Claim 7 or a pharmaceutically acceptable salt
 20 thereof, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-
 25 tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl,
 chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a

higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



5 R_2 is selected from the group consisting of hydrido, methyl, ethyl, phenyl, 4-trifluoromethylphenyl, trifluoromethoxymethyl, 1,1,2,2-tetrafluoroethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

10 R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, trifluoromethyl, difluoromethyl, and chlorodifluoromethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

15 Y is a covalent single bond;
Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

- R_5 and R_{10} are independently selected from the group consisting of
- benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy,
 3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy,
 3-chlorobenzyloxy, 2-chlorophenoxy, 4-chloro-3-ethylphenoxy,
 5 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy,
 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy,
 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy,
 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy,
 cyclobutoxy, cyclobutyl, cyclohexylmethoxy, cyclopentoxy, cyclopentyl,
 10 cyclopentylcarbonyl, cyclopropylmethoxy, 2,3-dichlorophenoxy,
 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl,
 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy,
 2,3-difluorobenzyloxy, 3,5-difluorobenzyloxy, difluoromethoxy,
 3,5-difluorophenoxy, 3,4-difluorophenyl, 2,3-difluorophenoxy,
 15 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy,
 3-dimethylaminophenoxy, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy,
 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 1,3-dioxolan-2-yl,
 3-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,
 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylbenzyl, 4-fluorobenzyloxy,
 20 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy,
 3-fluoro-5-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy,
 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy, 2-furyl, 3-furyl,
 25 heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl,
 2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl,
 4-isoxazolyl, 5-isoxazolyl, isopropoxy,
 3-isopropylbenzyloxy, 3-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl,
 30 3-methoxybenzyl, 4-methoxyphenylamino, 3-methylbenzyloxy,

- 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl, pentafluoroethyl,
 5 pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy,
 phenylamino, 1-phenylethoxy, 4-propylphenoxy, 4-propoxyphenoxy,
 thiophen-3-yl, tert-butoxy, 3-tert-butylphenoxy, 4-tert-butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 10 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl,
 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
 15 3-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy,
 3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy,
 20 3-pentafluoroethylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy,
 3-trifluoromethylthiophenoxy, 3-trifluoromethylthiobenzyloxy, and
 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and
 25 trifluoromethyl;

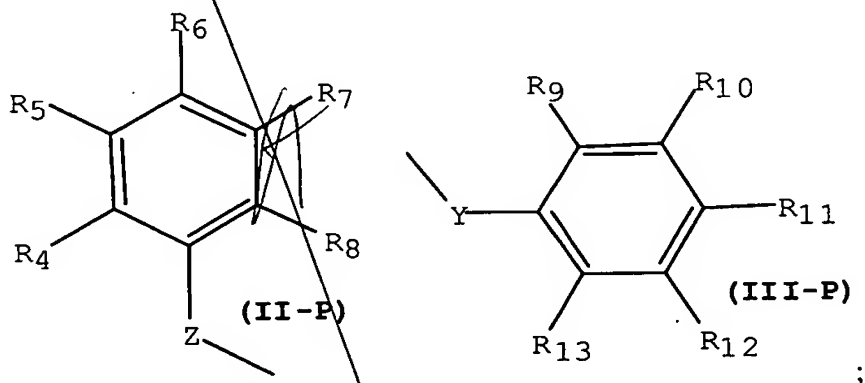
R_7 and R_{12} are independently selected from the group consisting of
 hydrido, fluoro, and trifluoromethyl.

9. The compound as recited in Claim 6 or a pharmaceutically acceptable salt thereof wherein;

n is the integer 1;

5 R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, methyl, ethyl, propyl, butyl, vinyl, phenyl, 4-trifluoromethylphenyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, ethyl, vinyl, trifluoromethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and

pentafluoroethyl with the provisos that $(\text{CHR}_3)_n\text{-N(Ap)Qp}$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is oxy;

5 Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 and R_{10} are independently selected from the group consisting of 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 10 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy, 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 15 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl, 20 cyclohexoxy, cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorobenzyloxy, 25 3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 3,5-difluorobenzyloxy, 4-difluoromethoxybenzyloxy, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy,

- 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy,
 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy,
 3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl,
 1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy,
 5 4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,
 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl,
 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy,
 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy,
 10 3-fluoro-5-trifluoromethylbenzyloxy, 4-fluoro-2-trifluoromethylbenzyloxy,
 4-fluoro-3-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy,
 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
 15 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
 3-iodobenzyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzyloxy,
 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl,
 20 5-isothiazolyl, 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
 3-methoxycarbonylprop-2-en-yloxy, 4-methoxyphenyl,
 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy,
 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 25 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl,
 pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy,
 phenylamino, 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy,
 30 propoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, *sec*-butyl,

4-*sec*-butylphenoxy, *tert*-butoxy, 3-*tert*-butylphenoxy, 4-*tert*-butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 2-(5,6,7,8-tetrahydronaphthoxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy,
 5 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl,
 3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy,
 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 10 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,
 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
 15 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, and
 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy,
 20 trifluoromethyl, and trifluoromethoxy;

R_7 and R_{12} are independently selected from the group consisting of
 hydrido, fluoro, and trifluoromethyl.

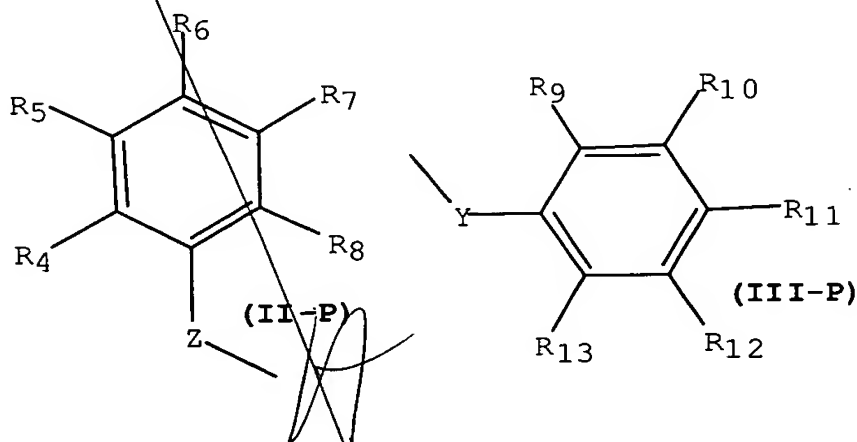
10. The compound as recited in Claim 9 or a pharmaceutically acceptable salt
 25 thereof, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and

5 $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, methyl, ethyl, phenyl, 4-trifluoromethylphenyl, trifluoromethoxymethyl, 1,1,2,2-tetrafluoroethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog

10 system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, trifluoromethyl, difluoromethyl, and

15 chlorodifluoromethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is oxy;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

- 5 R_5 and R_{10} are independently selected from the group consisting of benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy, 3-chlorobenzyloxy, 2-chlorophenoxy, 4-chloro-3-ethylphenoxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy,
- 10 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, cyclobutoxy, cyclobutyl, cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl, cyclopropylmethoxy, 2,3-dichlorophenoxy,
- 15 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 3,5-difluorobenzyloxy, difluoromethoxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy,
- 20 3-dimethylaminophenoxy, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 1,3-dioxolan-2-yl, 3-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylbenzyl, 4-fluorobenzyloxy, 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
- 25 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy, 2-furyl, 3-furyl, heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl,

- 2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl,
4-isoxazolyl, 5-isoxazolyl, isopropoxy,
3-isopropylbenzyloxy, 3-isopropylphenoxy, isopropylthio,
4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl,
5 3-methoxybenzyl, 4-methoxyphenylamino, 3-methylbenzyloxy,
4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
4-methylthiophenoxy, 2-naphthyl, 2-nitrophenoxy, 4-nitrophenoxy,
3-nitrophenyl, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl, pentafluoroethyl,
10 pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy,
phenylamino, 1-phenylethoxy, 4-propylphenoxy, 4-propoxyphenoxy,
thiophen-3-yl, tert-butoxy, 3-tert-butylphenoxy, 4-tert-butylphenoxy,
1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
15 2-(5,6,7,8-tetrahydronaphthyl), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
thiophen-2-yl, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl,
3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
20 3-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
3-trifluoromethylphenyl, 2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy,
3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy,
25 3-pentafluoroethylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy,
3-trifluoromethylthiophenoxy, 3-trifluoromethylthiobenzyloxy, and
trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and
30 trifluoromethyl;

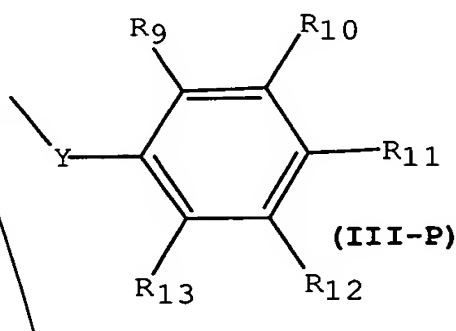
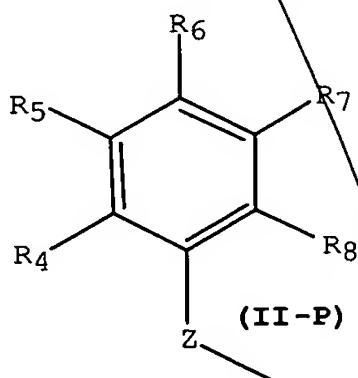
R_7 and R_{12} are independently selected from the group consisting of hydrido, fluoro, and trifluoromethyl.

11. The compound as recited in Claim 6 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, methyl, ethyl, propyl, butyl, vinyl, phenyl, 4-trifluoromethylphenyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, ethyl, vinyl, trifluoromethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-

- 5 Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of methylene and ethylene;
Z is a covalent single bond;

- $R_4, R_8, R_9,$ and R_{13} are independently selected from the group
10 consisting of hydrido and fluoro;

- R_5 and R_{10} are independently selected from the group consisting of 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy,
15 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy,
20 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl, cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl, cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy,
25 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorobenzyloxy,

- 3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, 3,5-difluorophenoxy,
 3,4-difluorophenyl, 3,5-difluorobenzyloxy, 4-difluoromethoxybenzyloxy,
 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy,
 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy,
 5 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy,
 3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl,
 1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy,
 4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,
 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl,
 10 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy,
 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy,
 3-fluoro-5-trifluoromethylbenzyloxy, 4-fluoro-2-trifluoromethylbenzyloxy,
 4-fluoro-3-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
 15 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy,
 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
 3-iodobenzyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
 20 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzyloxy,
 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl,
 5-isothiazolyl, 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
 3-methoxycarbonylprop-2-enyloxy, 4-methoxyphenyl,
 25 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy,
 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl,
 30 pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,

- 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy, propoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, *sec*-butyl, 4-*sec*-butylphenoxy, *tert*-butoxy, 3-*tert*-butylphenoxy, 4-*tert*-butylphenoxy,
- 5 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl, 2-(5,6,7,8-tetrahydronaphthyl)oxy, thiazol-2-yl, thiazol-4-yl, thiazol-5-yl, thiophen-2-yl, 2,3,5-trifluorobenzoyloxy, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy, 3-trifluoromethoxybenzoyloxy, 4-trifluoromethoxybenzoyloxy,
- 10 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl, 3-trifluoromethylbenzoyloxy, 4-trifluoromethylbenzoyloxy, 2,4-bis-trifluoromethylbenzoyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl, 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzoyloxy, 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
- 15 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzoyloxy, 4-trifluoromethylthiobenzoyloxy, 2,3,4-trifluorophenoxy, 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, and
- 20 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, trifluoromethyl, and trifluoromethoxy;

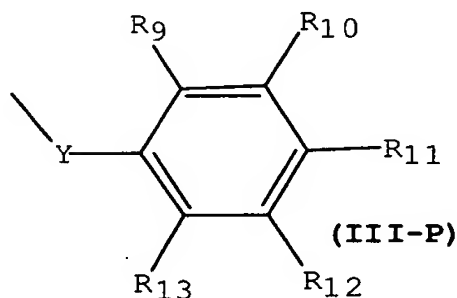
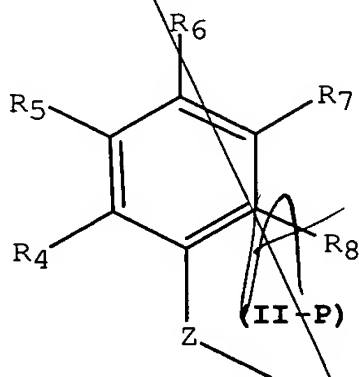
- R_7 and R_{12} are independently selected from the group consisting of
- 25 hydrido, fluoro, and trifluoromethyl.

12. The compound as recited in Claim 11 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, methyl, ethyl, phenyl, 4-trifluoromethylphenyl, trifluoromethoxymethyl, 1,1,2,2-tetrafluoroethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, trifluoromethyl, difluoromethyl, and chlorodifluoromethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower

Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher

Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is methylene;

Z is a covalent single bond;

5 R_4, R_8, R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 and R_{10} are independently selected from the group consisting of benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy, 10 3-chlorobenzyloxy, 2-chlorophenoxy, 4-chloro-3-ethylphenoxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 15 cyclobutoxy, cyclobutyl, cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl, cyclopropylmethoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 3,5-difluorobenzyloxy, difluoromethoxy, 20 3,5-difluorophenoxy, 3,4-difluorophenyl, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 1,3-dioxolan-2-yl, 3-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy, 25 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylbenzyl, 4-fluorobenzyloxy, 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy,

- 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzoyloxy,
 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy, 2-furyl, 3-furyl,
 heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl,
 2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl,
 5 4-isoxazolyl, 5-isoxazolyl, isopropoxy,
 3-isopropylbenzyloxy, 3-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl,
 3-methoxybenzyl, 4-methoxyphenylamino, 3-methylbenzyloxy,
 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 10 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl, pentafluoroethyl,
 pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy,
 15 phenylamino, 1-phenylethoxy, 4-propylphenoxy, 4-propoxyphenoxy,
 thiophen-3-yl, tert -butoxy, 3-tert -butylphenoxy, 4-tert -butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl,
 20 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
 3-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 25 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy,
 3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy,
 3-pentafluoroethylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy,
 3-trifluoromethylthiophenoxy, 3-trifluoromethylthiobenzyloxy, and
 30 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and trifluoromethyl;

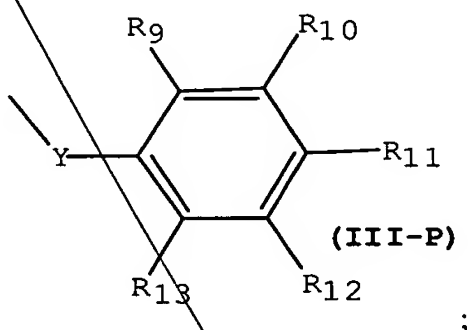
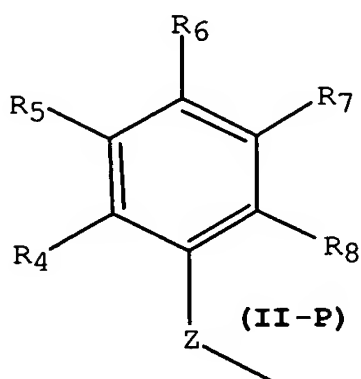
R_7 and R_{12} are independently selected from the group consisting of hydrido, fluoro, and trifluoromethyl.

13. The compound as recited in Claim 6 or a pharmaceutically acceptable salt, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is haloalkyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(\text{CHR}_3)_n\text{-N}(\text{Ap})\text{Qp}$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, alkyl, haloalkyl, aryl, and haloalkoxy with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, alkyl, and haloalkyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is a covalent single bond;

Z is covalent single bond;

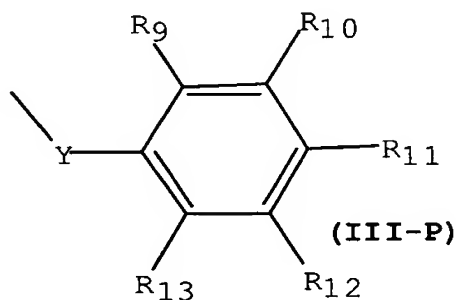
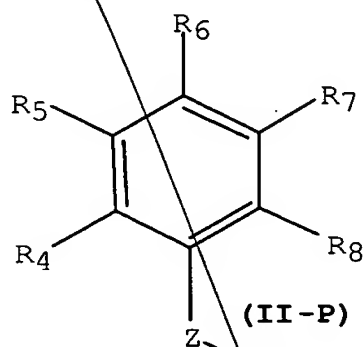
R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, heterocycloxy, aralkylaryl, heteroaryloxyalkyl, heteroarylthio, and heteroarylsulfonyl.

14. The compound as recited in Claim 13 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

R_1 is selected from the group consisting of trifluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



5

R_{16} is hydrido;

R_2 is selected from the group consisting of hydrido and phenyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

10 R_3 is selected from the group consisting of hydrido, methyl, trifluoromethyl, and difluoromethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is a covalent single bond;

15

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_3 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy, 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of cyclopentyl, 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl, isobutyl, isopropoxy, pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of fluoro and hydrido;

R_7 and R_{12} are independently selected from the group consisting of hydrido and fluoro.

15. The compound as recited in Claim 14 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

R_1 is selected from the group consisting of trifluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_{16} is hydrido;

R_2 is hydrido;

R_3 is hydrido;

Y is a covalent single bond;

5 Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 10 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy), 15 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of 1,1,2,2-tetrafluoroethoxy, pentafluoroethyl, and trifluoromethyl;

R_6 and R_{11} are independently selected from the group consisting of 20 fluoro and hydrido;

R_7 and R_{12} are independently selected from the group consisting of hydrido and fluoro.

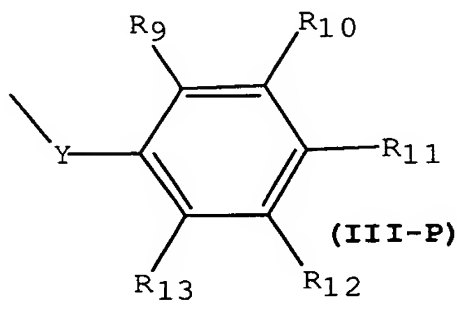
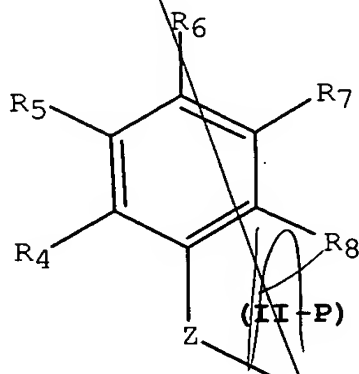
16. The compound as recited in Claim 6 or a pharmaceutically acceptable 25 salt, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is haloalkyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$

5 wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, alkyl, haloalkyl, aryl, and haloalkoxy with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, alkyl, and haloalkyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is oxy;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, heterocyclyloxy, aralkylaryl, heteroaryloxyalkyl, heteroarylthio, and heteroarylsulfonyl.

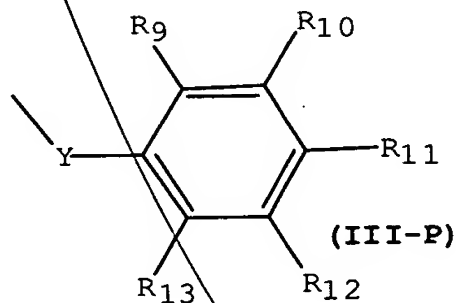
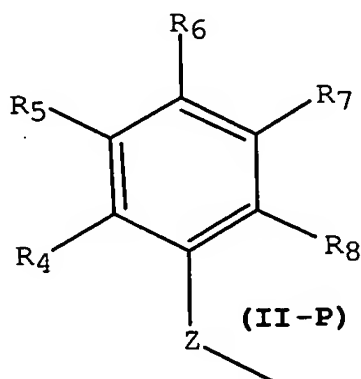
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17. The compound as recited in Claim 16 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

15

R_1 is selected from the group consisting of trifluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



;

R_{16} is hydrido;

R_2 is selected from the group consisting of hydrido and phenyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

5 R_3 is selected from the group consisting of hydrido, methyl, trifluoromethyl, and difluoromethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is oxy;

10 Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 15 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthoxy), 20 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of cyclopentyl, 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl, isobutyl,

isopropoxy, pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of fluoro and hydrido;

5 R_7 and R_{12} are independently selected from the group consisting of hydrido and fluoro.

18. The compound as recited in Claim 17 or a pharmaceutically acceptable salt thereof, wherein;

10

n is the integer 1;

R_1 is selected from the group consisting of trifluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_{16} is hydrido;

15

R_2 is hydrido;

R_3 is hydrido;

Y is oxy;

Z is a covalent single bond;

20 R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy,

4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyloxy), 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy,
 5 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of 1,1,2,2-tetrafluoroethoxy, pentafluoroethyl, and trifluoromethyl;

R_6 and R_{11} are independently selected from the group consisting of fluoro and hydrido;

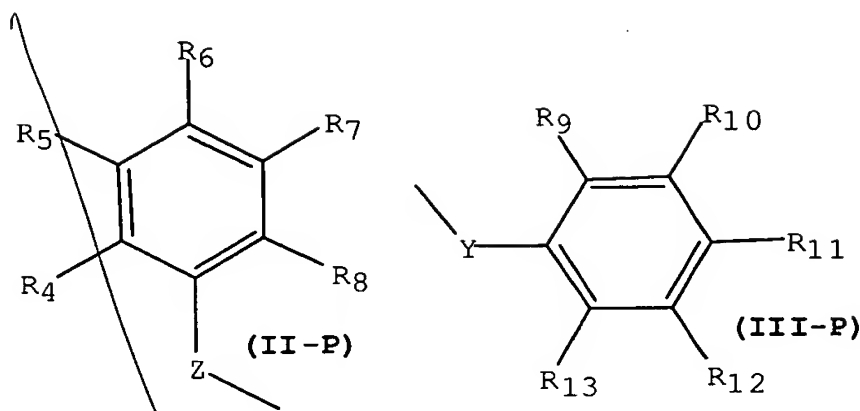
10 R_7 and R_{12} are independently selected from the group consisting of hydrido and fluoro.

15 19. The compound as recited in Claim 6 or a pharmaceutically acceptable salt, wherein;

n is the integer 1;

R_{16} is hydrido;

20 R_1 is haloalkyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(Ap)Qp$ wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_2 is selected from the group consisting of hydrido, alkyl, haloalkyl, aryl, and haloalkoxy with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, alkyl, and haloalkyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is methylene;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

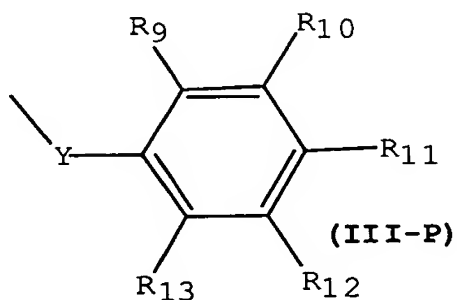
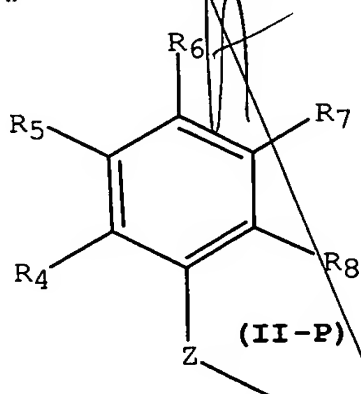
R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl,

heteroaralkoxy, heterocycloxy, aralkylaryl, heteroaryloxyalkyl,
heteroarylthio, and heteroarylsulfonyl.

20. The compound as recited in Claim 19 or a pharmaceutically acceptable
5 salt thereof, wherein;

n is the integer 1;

R_1 is selected from the group consisting of trifluoromethyl,
chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a
10 higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and
(CHR_3) _{n} -N(Ap)Qp wherein Ap is Formula (II-P) and Qp is Formula (III-P);



R_{16} is hydrido;

15 R_2 is selected from the group consisting of hydrido and phenyl with
the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both
 R_1 and (CHR_3) _{n} -N(Ap)Qp;

R_3 is selected from the group consisting of hydrido, methyl, trifluoromethyl, and difluoromethyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

5 Y is methylene;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

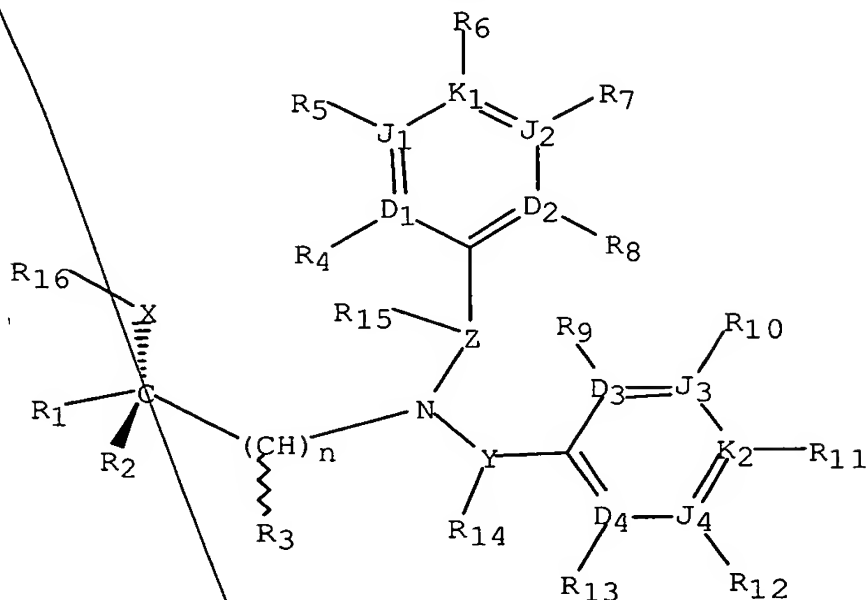
R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 10 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthoxy), 15 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of cyclopentyl, 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl, isobutyl, 20 isopropoxy, pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of fluoro and hydrido;

R_7 and R_{12} are independently selected from the group consisting of 25 hydrido and fluoro.

21. A compound as recited in Claim 2 having the formula:



5 or a pharmaceutically acceptable salt thereof, wherein;

D_1, D_2, J_1, J_2 and K_1 are each carbon with the proviso that at least one of D_3, D_4, J_3, J_4 and K_2 is selected from the group consisting of O, S, and N, wherein D_3, D_4, J_3, J_4 and K_2 are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that no more than one of D_3, D_4, J_3, J_4 and K_2 is a covalent bond, no more than one of D_3, D_4, J_3, J_4 and K_2 is O, no more than one of D_3, D_4, J_3, J_4 and K_2 is S, one of D_3, D_4, J_3, J_4 and K_2 must be a covalent bond when two of D_3, D_4, J_3, J_4 and K_2 are O and S, and no more than four of D_3, D_4, J_3, J_4 and K_2 are N;

D_1, D_2, J_1, J_2 and K_1 are independently selected from the group consisting of C, O, S, N and covalent bond with the provisos that $D_3, D_4, J_3,$

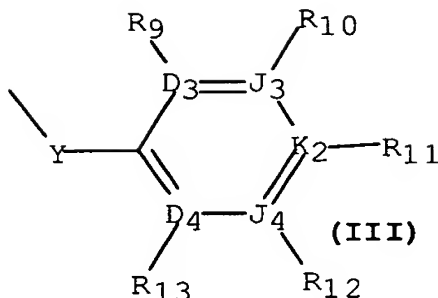
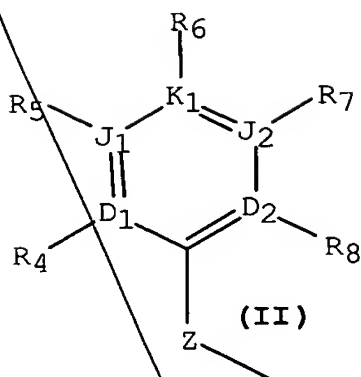
J_4 and K_2 are each carbon and at least one of D_1 , D_2 , J_1 , J_2 and K_1 is selected from the group consisting of O, S, and N wherein, when D_1 , D_2 , J_1 , J_2 and K_1 are selected from the group consisting of C, O, S, covalent bond, and N, no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is a covalent bond, no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is O, no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is S, one of D_1 , D_2 , J_1 , J_2 and K_1 must be a covalent bond when two of D_1 , D_2 , J_1 , J_2 and K_1 are O and S, and no more than four of D_1 , D_2 , J_1 , J_2 and K_1 are N;

n is an integer selected from 1 and 2;

X is oxy;

R_{16} is selected from the group consisting of hydrido and a spacer selected from the group consisting of a covalent single bond and a linear spacer moiety having a chain length of 1 to 4 atoms linked to the point of bonding of any aromatic substituent selected from the group consisting of R_4 , R_8 , R_9 , and R_{13} to form a heterocyclyl ring having from 5 through 10 contiguous members;

R_1 is selected from the group consisting of haloalkyl and haloalkoxymethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(A)Q$ wherein A is Formula (II) and Q is Formula (III);



R_2 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, haloalkoxy, haloalkoxyalkyl, perhaloaryl, perhaloaralkyl, perhaloaryloxyalkyl, and heteroaryl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(Ap)Qp$;

R_3 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, and haloalkoxyalkyl with the provisos that $(CHR_3)_n-N(Ap)Qp$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of a covalent single bond, $(CH_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH_2)_j-O-(CH_2)_k$ wherein j and k are integers independently selected from 0 and 1;

Z is selected from the group consisting of covalent single bond, $(CH_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH_2)_j-O-(CH_2)_k$ wherein j and k are integers independently selected from 0 and 1;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido, halo, haloalkyl, and alkyl ;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, carboxy, heteroaralkylthio, heteroaralkoxy, cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy, heterocyclyloxy, aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocyclyl, perhaloaralkyl,

5 aralkylsulfonyl, aralkylsulfonylalkyl, aralkylsulfinyl, aralkylsulfinylalkyl, halocycloalkyl, halocycloalkenyl, cycloalkylsulfinyl, cycloalkylsulfinylalkyl, cycloalkylsulfonyl, cycloalkylsulfonylalkyl, heteroaryl amino, N-heteroaryl amino, N-alkyl amino, heteroaryl amino alkyl, haloalkylthio, alkanoyloxy, alkoxy, alkoxyalkyl, haloalkoxyalkyl, heteroaralkoxy,

10 cycloalkoxy, cycloalkenyloxy, cycloalkoxyalkyl, cycloalkylalkoxy, cycloalkenyloxyalkyl, cycloalkylenedioxy, halocycloalkoxy, halocycloalkoxyalkyl, halocycloalkenyloxy, halocycloalkenyloxyalkyl, hydroxy, amino, thio, nitro, lower alkyl amino, alkylthio, alkylthioalkyl, aryl amino, aralkyl amino, arylthio, arylthioalkyl, heteroaralkoxyalkyl,

15 alkylsulfinyl, alkylsulfinylalkyl, arylsulfinylalkyl, arylsulfonylalkyl, heteroaryl sulfinylalkyl, heteroaryl sulfonylalkyl, alkylsulfonyl, alkylsulfonylalkyl, haloalkylsulfinylalkyl, haloalkylsulfonylalkyl, alkylsulfonamido, alkylaminosulfonyl, amidosulfonyl, monoalkyl amidosulfonyl, dialkyl amidosulfonyl, monoaryl amidosulfonyl,

20 arylsulfonamido, diarylamidosulfonyl, monoalkyl monoaryl amidosulfonyl, arylsulfinyl, arylsulfonyl, heteroarylthio, heteroaryl sulfinyl, heteroaryl sulfonyl, heterocyclylsulfonyl, heterocyclylthio, alkanoyl, alkenoyl, aroyl, heteroaroyl, aralkanoyl, heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl, alkynyl, alkenyloxy, alkenyloxyalkyl, alkylenedioxy, haloalkylenedioxy,

25 cycloalkyl, cycloalkylalkanoyl, cycloalkenyl, lower cycloalkylalkyl, lower cycloalkenylalkyl, halo, haloalkyl, haloalkenyl, haloalkoxy, hydroxyhaloalkyl, hydroxyaralkyl, hydroxyalkyl, hydroxyheteroaralkyl, haloalkoxyalkyl, aryl, heteroaralkynyl, aryloxy, aralkoxy, aryloxyalkyl, saturated heterocyclyl, partially saturated heterocyclyl, heteroaryl,

30 heteroaryloxy, heteroaryloxyalkyl, heteroaralkyl, arylalkenyl,

heteroarylalkenyl, carboxyalkyl, carboalkoxy, alkoxy-carboxamido, alkylamidocarbonylamido, arylamidocarbonylamido, carboalkoxyalkyl, carboalkoxyalkenyl, carboaralkoxy, carboxamido, carboxamidoalkyl, cyano, carbohaloalkoxy, phosphono, phosphonoalkyl, diaralkoxyphosphono, and
 5 diaralkoxyphosphonoalkyl;

R_4 and R_5 , R_5 and R_6 , R_6 and R_7 , R_7 and R_8 , R_9 and R_{10} , R_{10} and R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} are independently selected to form spacer pairs wherein a spacer pair is taken together to form a linear moiety having from 3 through 6 contiguous atoms connecting the points of bonding of said
 10 spacer pair members to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 contiguous members, a partially saturated heterocyclic ring having 5 through 8 contiguous members, a heteroaryl ring having 5 through 6 contiguous members, and an aryl with the provisos that no more than one of the group consisting of spacer pairs R_4 and
 15 R_5 , R_5 and R_6 , R_6 and R_7 , and R_7 and R_8 , is used at the same time and that no more than one of the group consisting of spacer pairs R_9 and R_{10} , R_{10} and R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} is used at the same time.

22. The compound as recited in Claim 21 or a pharmaceutically acceptable
 20 salt thereof, wherein;

D_1 , D_2 , J_1 , J_2 and K_1 are each carbon with the proviso that at least one of D_3 , D_4 , J_3 , J_4 and K_2 is selected from the group consisting of O, S, and N, wherein D_3 , D_4 , J_3 , J_4 and K_2 are independently selected from the group
 25 consisting of C, N, O, S and covalent bond with the provisos that no more

than one of D_3, D_4, J_3, J_4 and K_2 is a covalent bond, no more than one of D_3, D_4, J_3, J_4 and K_2 is O, no more than one of D_3, D_4, J_3, J_4 and K_2 is S, one of D_3, D_4, J_3, J_4 and K_2 must be a covalent bond when two of D_3, D_4, J_3, J_4 and K_2 are O and S, and no more than four of D_3, D_4, J_3, J_4 and K_2 are N;

5 D_1, D_2, J_1, J_2 and K_1 are independently selected from the group consisting of C, O, S, N and covalent bond with the provisos that D_3, D_4, J_3, J_4 and K_2 are each carbon and at least one of D_1, D_2, J_1, J_2 and K_1 is selected from the group consisting of O, S, and N wherein, when D_1, D_2, J_1, J_2 and K_1 are selected from the group consisting of C, O, S, covalent bond, and N, no
10 more than one of D_1, D_2, J_1, J_2 and K_1 is a covalent bond, no more than one of D_1, D_2, J_1, J_2 and K_1 is O, no more than one of D_1, D_2, J_1, J_2 and K_1 is S, one of D_1, D_2, J_1, J_2 and K_1 must be a covalent bond when two of D_1, D_2, J_1, J_2 and K_1 are O and S, and no more than four of D_1, D_2, J_1, J_2 and K_1 are N;

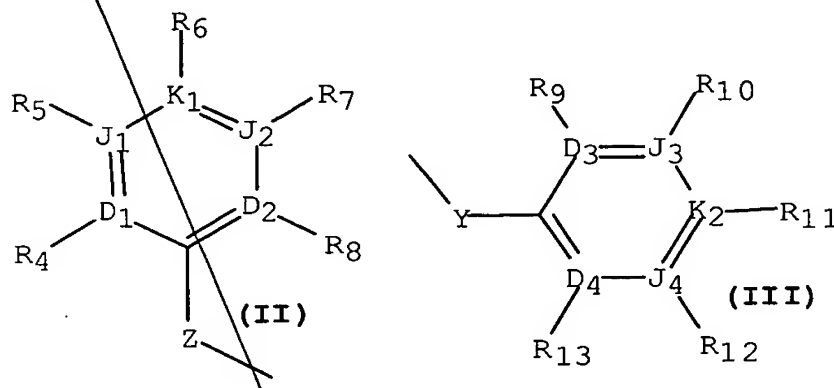
n is the integer 1;

15 X is oxy;

R_{16} is taken together with R_4, R_8, R_9 , or R_{13} to form a spacer selected from the group consisting of a covalent single bond, CH_2 , $CH(CH_3)$, CF_2 , $C(O)$, $C(S)$, and SO_2 ;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl,
20 chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a

higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(A)Q$ wherein A is Formula (II) and Q is Formula (III);



R_2 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, vinyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

R_3 is selected from the group consisting of hydrido, methyl, ethyl, vinyl, phenyl, 4-trifluoromethylphenyl, , trifluoromethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of covalent single bond, oxy, methyleneoxy, methylene, and ethylene;

Z is selected from the group consisting of covalent single bond, oxy, methyleneoxy, methylene, and ethylene;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 and R_{10} are independently selected from the group consisting of 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy, 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl, cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl, cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorobenzyloxy, 3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 3,5-difluorobenzyloxy, 4-difluoromethoxybenzyloxy, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl, 1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy, 4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl, 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy,

- 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy,
 3-fluoro-5-trifluoromethylbenzyloxy, 4-fluoro-2-trifluoromethylbenzyloxy,
 4-fluoro-3-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
 5 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy,
 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
 3-iodobenzyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
 10 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzyloxy,
 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl,
 5-isothiazolyl, 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
 3-methoxycarbonylprop-2-en-yloxy, 4-methoxyphenyl,
 15 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy,
 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 4-methylthiophenoxy, 2-naphthyl, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl,
 20 pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy,
 phenylamino, 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy,
 propoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, *sec*-butyl,
 4-*sec*-butylphenoxy, *tert*-butoxy, 3-*tert*-butylphenoxy, 4-*tert*-butylphenoxy,
 25 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 2-(5,6,7,8-tetrahydronaphthyl), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy,
 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 30 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl,

3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy,
 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 5 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,
 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, and
 10 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, difluoromethoxy, trifluoromethyl, trifluoromethoxy,
 pentafluoroethyl, and 1,1,2,2-tetrafluoroethoxy;

R_7 and R_{12} are independently selected from the group consisting of
 15 hydrido, fluoro, and trifluoromethyl.

23. The compound as recited in Claim 21 or a pharmaceutically acceptable
 salt thereof, wherein:

20 D_1, D_2, J_1, J_2 and K_1 are each carbon with the proviso that at least one
 of D_3, D_4, J_3, J_4 and K_2 is selected from the group consisting of O, S, and N,
 wherein D_3, D_4, J_3, J_4 and K_2 are independently selected from the group
 consisting of C, N, O, S and covalent bond with the provisos that no more
 than one of D_3, D_4, J_3, J_4 and K_2 is a covalent bond, no more than one of $D_3,$
 25 D_4, J_3, J_4 and K_2 is O, no more than one of D_3, D_4, J_3, J_4 and K_2 is S, one of

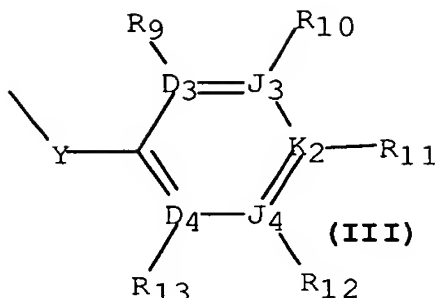
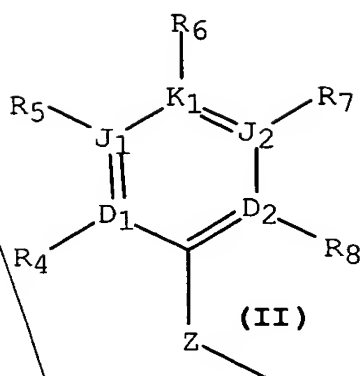
D_3, D_4, J_3, J_4 and K_2 must be a covalent bond when two of D_3, D_4, J_3, J_4 and K_2 are O and S, and no more than four of D_3, D_4, J_3, J_4 and K_2 are N;

D_1, D_2, J_1, J_2 and K_1 are selected from the group consisting of C, O, S, N and covalent bond with the provisos that D_3, D_4, J_3, J_4 and K_2 are each carbon and at least one of D_1, D_2, J_1, J_2 and K_1 is selected from the group consisting of O, S, and N wherein, when D_1, D_2, J_1, J_2 and K_1 are selected from the group consisting of C, O, S, covalent bond, and N, no more than one of D_1, D_2, J_1, J_2 and K_1 is a covalent bond, no more than one of D_1, D_2, J_1, J_2 and K_1 is O, no more than one of D_1, D_2, J_1, J_2 and K_1 is S, one of D_1, D_2, J_1, J_2 and K_1 must be a covalent bond when two of D_1, D_2, J_1, J_2 and K_1 are O and S, and no more than four of D_1, D_2, J_1, J_2 and K_1 are N;

n is an integer selected from 1 and 2;

X is oxy;

R_1 is selected from the group consisting of haloalkyl and haloalkoxymethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(A)Q$ wherein A is Formula (II) and Q is Formula (III);



R_{16} is hydrido;

R_2 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, and haloalkoxyalkyl with the proviso that R_2 has a lower

5 Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

R_3 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, and haloalkoxyalkyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

10 Y is selected from the group consisting of a covalent single bond, oxy and C1-C2 alkylene;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

15 R_4 , R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl,

heteroaralkoxy, heterocycloxy, aralkylaryl, heteroaryloxyalkyl,
heteroarylthio, and heteroarylsulfonyl.

- 5 24. The compound as recited in Claim 23 and pharmaceutically acceptable salts, wherein;

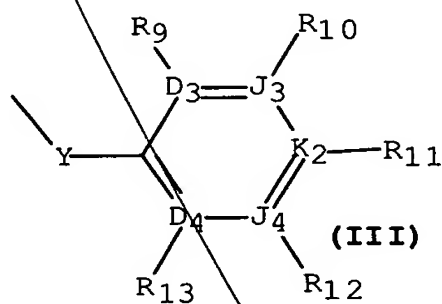
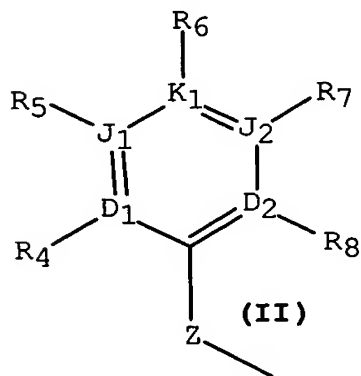
n is the integer 1;

X is oxy;

10 R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and

15 $(CHR_3)_n-N(A)Q$ wherein A is Formula (II) and Q is Formula (III);



R_2 is selected from the group consisting of hydrido, methyl, ethyl, propyl, butyl, vinyl, phenyl, 4-trifluoromethylphenyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, and

2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, ethyl, vinyl, trifluoromethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is selected from the group consisting of a single covalent bond, methylene, ethylene, and oxy;

Z is covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 and R_{10} are independently selected from the group consisting of 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy, 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy,

- 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl,
cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl,
cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy,
2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl,
5 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy,
3,4-difluorophenoxy, 2,3-difluorobenzoyloxy, 2,4-difluorobenzoyloxy,
3,4-difluorobenzoyloxy, 2,5-difluorobenzoyloxy, 3,5-difluorophenoxy,
3,4-difluorophenyl, 3,5-difluorobenzoyloxy, 4-difluoromethoxybenzoyloxy,
2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy,
10 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy,
3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzoyloxy,
3,5-dimethylbenzoyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl,
1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy,
4-ethylbenzoyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,
15 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl,
4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzoyloxy,
2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzoyloxy,
3-fluoro-5-trifluoromethylbenzoyloxy, 4-fluoro-2-trifluoromethylbenzoyloxy,
20 4-fluoro-3-trifluoromethylbenzoyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzoyloxy,
4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
25 3-iodobenzoyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzoyloxy,
3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,
4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl,
5-isothiazolyl, 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
30 3-methoxycarbonylprop-2-enyloxy, 4-methoxyphenyl,

- 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy, 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy, 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy, 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy,
- 5 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy, propoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, *sec*-butyl,
- 10 4-*sec*-butylphenoxy, *tert*-butoxy, 3-*tert*-butylphenoxy, 4-*tert*-butylphenoxy, 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl, 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl, thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
- 15 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenyl, 4-trifluoromethoxyphenoxy, trifluoromethyl, 3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy, 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl, 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
- 20 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy, 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy, 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy, 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
- 25 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, trifluoromethyl, and trifluoromethoxy;

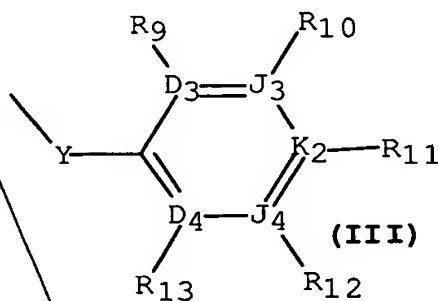
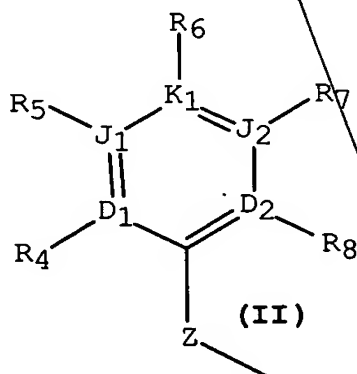
R_7 and R_{12} are independently selected from the group consisting of hydrido, fluoro, and trifluoromethyl.

- 5 25. The compound as recited in Claim 24 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

X is oxy;

- 10 R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(A)Q$ wherein A is Formula (II) and Q is Formula (III);



15

R_{16} is hydrido;

R_2 is selected from the group consisting of hydrido, methyl, ethyl, phenyl, 4-trifluoromethylphenyl, trifluoromethoxymethyl,

1,1,2,2-tetrafluoroethoxymethyl, difluoromethyl, and 2,2,3,3,3-pentafluoropropyl with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

R_3 is selected from the group consisting of hydrido, phenyl, 4-trifluoromethylphenyl, methyl, trifluoromethyl, difluoromethyl, and chlorodifluoromethyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is methylene;

10 Z is covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 and R_{10} are independently selected from the group consisting of benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy, 3-chlorobenzyloxy, 2-chlorophenoxy, 4-chloro-3-ethylphenoxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, cyclobutoxy, cyclobutyl, cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl, cyclopropylmethoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 3,5-difluorobenzyloxy, difluoromethoxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 2,3-difluorophenoxy,

- 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy,
 3-dimethylaminophenoxy, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy,
 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 1,3-dioxolan-2-yl,
 3-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,
 5 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylbenzyl, 4-fluorobenzyloxy,
 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy,
 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy,
 3-fluoro-5-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy,
 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy,
 10 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy, 2-furyl, 3-furyl,
 heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl,
 2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl,
 4-isoxazolyl, 5-isoxazolyl, isopropoxy, 3-isopropylbenzyloxy,
 3-isopropylphenoxy, isopropylthio, 4-isopropyl-3-methylphenoxy,
 15 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl, 3-methoxybenzyl,
 4-methoxyphenylamino, 3-methylbenzyloxy, 4-methylbenzyloxy,
 3-methylphenoxy, 3-methyl-4-methylthiophenoxy, 4-methylphenoxy,
 1-methylpropoxy, 2-methylpyrid-5-yloxy, 4-methylthiophenoxy,
 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy, 3-nitrophenyl, 2-oxazolyl,
 20 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-
 pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl,
 phenoxy, phenylamino, 1-phenylethoxy, 4-propylphenoxy,
 4-propoxyphenoxy, thiophen-3-yl, tert-butoxy, 3-tert-butylphenoxy,
 4-tert-butylphenoxy, 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 25 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl,
 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
 30 3-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,

3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy,
 3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy,
 5 3-pentafluoroethylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy,
 3-trifluoromethylthiophenoxy, 3-trifluoromethylthiobenzyloxy, and
 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and
 10 trifluoromethyl;

R_7 and R_{12} are independently selected from the group consisting of
 hydrido, fluoro, and trifluoromethyl.

26. The compound as recited in Claim 23 or a pharmaceutically acceptable
 15 salt, wherein;

D_1 , D_2 , J_1 , J_2 and K_1 are each carbon;

D_3 , D_4 , J_3 , J_4 and K_2 are independently selected from the group
 consisting of C, N, O, S and covalent bond with the provisos that no more
 than one of D_3 , D_4 , J_3 , J_4 and K_2 is a covalent bond, no more than one of D_3 ,
 20 D_4 , J_3 , J_4 and K_2 is O, no more than one of D_3 , D_4 , J_3 , J_4 and K_2 is S, one of
 D_3 , D_4 , J_3 , J_4 and K_2 must be a covalent bond when two of D_3 , D_4 , J_3 , J_4 and
 K_2 are O and S, no more than four of D_3 , D_4 , J_3 , J_4 and K_2 are N, and one of
 D_3 , D_4 , J_3 , J_4 and K_2 is selected from the group consisting of O, S, and N;

25 n is the integer 1;

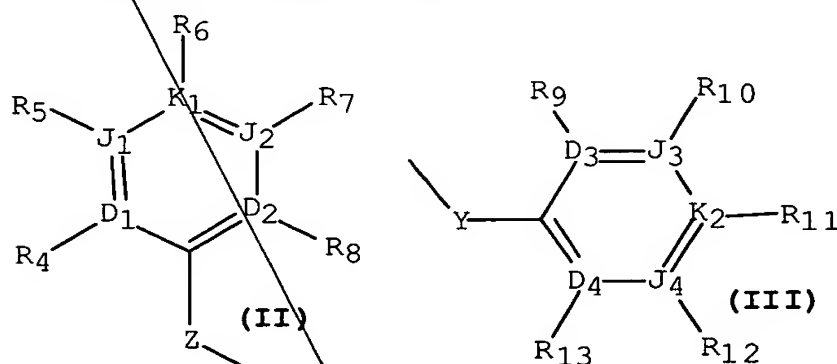
X is oxy;

R_{16} is hydrido;

R_1 is haloalkyl with the proviso that R_1 has a higher Cahn-Ingold-

Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(A)Q$

5 wherein A is Formula (II) and Q is Formula (III);



R_2 is selected from the group consisting of hydrido, alkyl, aryl, haloalkyl, and haloalkoxy with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

10 R_3 is selected from the group consisting of hydrido, alkyl, and haloalkyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is a C1-C2 alkylene;

15 Z is covalent single bond;

R_{14} is hydrido;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, and heteroaryloxyalkyl.

27. The compound as recited in Claim 23 or a pharmaceutically acceptable salt, wherein;

D_3 , D_4 , J_3 , J_4 and K_2 are each carbon;

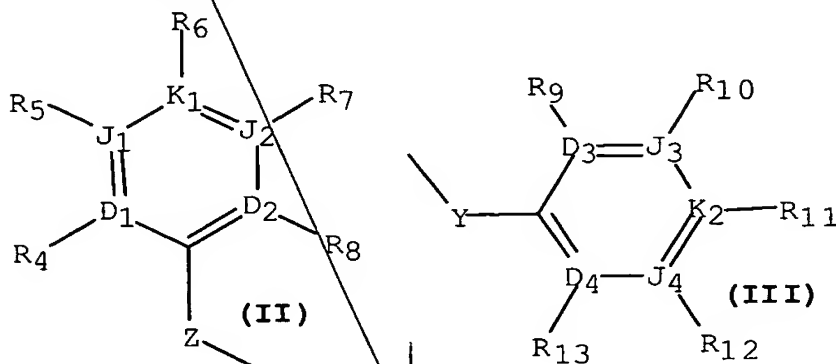
D_1 , D_2 , J_1 , J_2 and K_1 are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is a covalent bond, no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is O, no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is S, one of D_1 , D_2 , J_1 , J_2 and K_1 must be a covalent bond when two of D_1 , D_2 , J_1 , J_2 and K_1 are O and S, no more than four of D_1 , D_2 , J_1 , J_2 and K_1 are N, and one of D_1 , D_2 , J_1 , J_2 and K_1 is selected from the group consisting of O, S, and N;

n is the integer 1;

X is oxy;

R_{16} is hydrido;

R_1 is haloalkyl with the proviso that R_1 has a higher Cahn-Ingold-Prelog stereochemical system ranking than both R_2 and $(CHR_3)_n-N(A)Q$ wherein A is Formula (II) and Q is Formula (III);



5 R_2 is selected from the group consisting of hydrido, alkyl, aryl, haloalkyl, and haloalkoxy with the proviso that R_2 has a lower Cahn-Ingold-Prelog system ranking than both R_1 and $(CHR_3)_n-N(A)Q$;

R_3 is selected from the group consisting of hydrido, alkyl, and haloalkyl with the provisos that $(CHR_3)_n-N(A)Q$ has a lower Cahn-Ingold-Prelog stereochemical system ranking than R_1 and a higher Cahn-Ingold-Prelog stereochemical system ranking than R_2 ;

Y is a C1-C2 alkylene;

Z is covalent single bond;

R_{14} is hydrido;

15 $R_4, R_8, R_9,$ and R_{13} are independently selected from the group consisting of hydrido and halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, and heteroaryloxyalkyl.

28. The compound as recited in any one of Claims 26 or 27 or a pharmaceutically acceptable salt thereof, wherein;

10

n is the integer 1;

X is oxy;

R_1 is selected from the group consisting of trifluoromethyl and pentafluoroethyl and chlorodifluoromethyl;

15

R_{16} is hydrido;

R_2 is hydrido;

R_3 is selected from the group consisting of hydrido, methyl, trifluoromethyl, and difluoromethyl;

Y is methylene;

20

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3,5-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy,

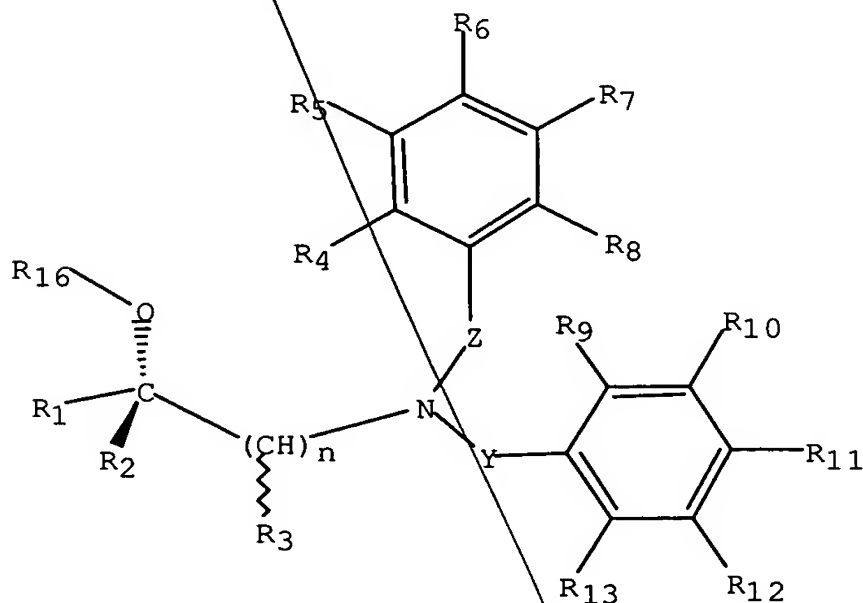
- 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthylloxy), 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of cyclopentyl, 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl, isobutyl, isopropoxy, pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

- R_6 and R_{11} are independently selected from the group consisting of fluoro and hydrido;

R_7 and R_{12} are independently selected from the group consisting of hydrido and fluoro.

29. A compound having the formula:



or a pharmaceutically acceptable salt thereof, wherein:

n is an integer selected from 1 and 2;

R_1 is selected from the group consisting of haloalkyl and haloalkoxyalkyl;

5 R_{16} is hydrido;

R_2 is hydrido;

R_3 is hydrido;

Y is selected from the group consisting of a covalent single bond and C1-C2 alkylene;

10 Z is selected from the group consisting of a covalent single bond and C1-C2 alkylene;

15 $R_4, R_8, R_9,$ and R_{13} are independently selected from the group consisting of hydrido and halo;

$R_5, R_6, R_7, R_{10}, R_{11},$ and R_{12} are independently selected from the group consisting of perhaloaryloxy, N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclalthio, hydroxyalkoxy, carboxamidoalkoxy, alkoxycarbonylalkoxy, alkoxycarbonylalkenyloxy, aralkanoylalkoxy, aralkenoyl, N-arylcarboxamidoalkoxy, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl, hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, heterocyclyloxy, aralkylaryl, heteroaryloxyalkyl, heteroarylthio, and heteroarylsulfonyl.

25

30. The compound as recited in Claim 29 or a pharmaceutically acceptable salt thereof wherein;

5

n is the integer 1;

R_{16} is hydrido;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

10

R_2 is hydrido;

R_3 is hydrido;

Y is selected from the group consisting of methylene, and ethylene;

15

Z is selected from the group consisting of covalent single bond and methylene;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 and R_{10} are independently selected from the group consisting of 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy, 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy,

20

25

- 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy,
2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl,
cyclohexoxy, cyclohexylmethoxy, cyclopentoxo, cyclopentyl,
cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy,
2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl,
3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy,
3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorobenzyloxy,
3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, 3,5-difluorophenoxy,
3,4-difluorophenyl, 3,5-difluorobenzyloxy, 4-difluoromethoxybenzyloxy,
2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy,
3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy,
3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy,
3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl,
1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy,
4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy,
3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl,
4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy,
2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy,
3-fluorophenoxy, 3-fluoro-2-nitrophenoxy,
2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy,
4-fluoro-2-trifluoromethylbenzyloxy, 4-fluoro-3-trifluoromethylbenzyloxy,
2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy,
2-fluorobenzyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl, 1,1,1,3,3,3-
hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy, 3-iodobenzyloxy,
isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl, 4-isoxazolyl, 5-isoxazolyl,
isopropoxy, isopropyl, 4-isopropylbenzyloxy, 3-isopropylphenoxy,
4-isopropylphenoxy, isopropylthio, 4-isopropyl-3-methylphenoxy,
3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl, 3-methoxybenzyl,
4-methoxycarbonylbutoxy, 3-methoxycarbonylprop-2-enyloxy,

- 4-methoxyphenyl, 3-methoxyphenylamino, 4-methoxyphenylamino,
 3-methylbenzyloxy, 4-methylbenzyloxy, 3-methylphenoxy,
 3-methyl-4-methylthiophenoxy, 4-methylphenoxy, 1-methylpropoxy,
 2-methylpyrid-5-yloxy, 4-methylthiophenoxy, 2-naphthyloxy,
 5 2-nitrophenoxy, 4-nitrophenoxy, 3-nitrophenyl, 4-nitrophenylthio, 2-
 oxazolyl, 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio,
 2,2,3,3,3-pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl,
 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy,
 phenylsulfonyl, 4-propanoylphenoxy, propoxy, 4-propylphenoxy,
 10 4-propoxyphenoxy, thiophen-3-yl, *sec*-butyl, 4-*sec*-butylphenoxy,
tert-butoxy, 3-*tert*-butylphenoxy, 4-*tert*-butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy;
 15 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl, 3-
 trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy,
 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 20 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,
 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
 25 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, and
 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy,
 30 trifluoromethyl, and trifluoromethoxy;

R_7 and R_{12} are independently selected from the group consisting of hydrido, fluoro, and trifluoromethyl.

- 5 31. The compound as recited in Claim 30 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

- 10 R_1 is selected from the group consisting of trifluoromethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_{16} is hydrido;

R_2 is hydrido;

R_3 is hydrido;

Y is methylene;

- 15 Z is covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

- 20 R_5 and R_{10} are independently selected from the group consisting of benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy, 3-chlorobenzyloxy, 2-chlorophenoxy, 4-chloro-3-ethylphenoxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy,

- 4-chlorophenylamino, 5-chloropyrid-3-yloxy, cyclobutoxy, cyclobutyl, cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl, cyclopropylmethoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl,
- 5 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzoyloxy, 3,5-difluorobenzoyloxy, difluoromethoxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,4-dimethylbenzoyloxy, 3,5-dimethylbenzoyloxy, 3,5-dimethylphenoxy,
- 10 3,4-dimethylphenoxy, 1,3-dioxolan-2-yl, 3-ethylbenzoyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylbenzyl, 4-fluorobenzoyloxy, 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzoyloxy, 3-fluoro-5-trifluoromethylbenzoyloxy,
- 15 2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzoyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy, 2-furyl, 3-furyl, heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl, 4-isoxazolyl, 5-isoxazolyl, isopropoxy, 3-isopropylbenzoyloxy,
- 20 3-isopropylphenoxy, isopropylthio, 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl, 3-methoxybenzyl, 4-methoxyphenylamino, 3-methylbenzoyloxy, 4-methylbenzoyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy, 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy, 4-methylthiophenoxy,
- 25 2-naphthylloxy, 2-nitrophenoxy, 4-nitrophenoxy, 3-nitrophenyl, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, tert -butoxy,
- 30 3-tert -butylphenoxy, 4-tert -butylphenoxy, 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl, 2-(5,6,7,8-tetrahydronaphthylloxy), thiazol-2-yl,

thiazol-4-yl, thiazol-5-yl, thiophen-2-yl, 2,2,2-trifluoroethoxy,
 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
 5 3-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy,
 3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy,
 10 3-pentafluoroethylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy,
 3-trifluoromethylthiophenoxy, 3-trifluoromethylthiobenzyloxy, and
 trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and
 15 trifluoromethyl;

R_7 and R_{12} are independently selected from the group consisting of
 hydrido, fluoro, and trifluoromethyl.

32. The compound as recited in Claim 29 or a pharmaceutically acceptable
 20 salt, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is haloalkyl;

25 R_2 is is hydrido;

R_3 is is hydrido;

Y is methylene;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

5 R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of perhaloaryloxy, N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclthio, hydroxyalkoxy, aralkanoylalkoxy, aralkenoyl, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl, hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, 10 arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, and heteroaryloxyalkyl.

33. The compound as recited in Claim 32 or a pharmaceutically 15 acceptable salt, wherein;

n is the integer 1;

R_{16} is hydrido;

R_1 is haloalkyl;

20 R_2 is is hydrido;

R_3 is is hydrido;

Y is methylene;

Z is a covalent single bond;

25 R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and halo;

R_5 is selected from the group consisting of perhaloaryloxy,
 N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclthio, hydroxyalkoxy,
 aralkanoylalkoxy, aralkenoyl, cycloalkylcarbonyl, cyanoalkoxy,
 heterocyclcarbonyl, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino,
 5 arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy,
 aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl,
 cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, and
 heteroaryloxyalkyl.

R_{10} is selected from the group consisting of haloalkyl, haloalkoxy,
 10 aryl, alkylthio, alkoxy, aralkyl, alkyl, cycloalkoxy, cycloalkylalkoxy,
 heteroaryl, cycloalkyl, haloalkylthio, and hydroxyhaloalkyl.

R_6 and R_{11} are independently selected from the group consisting of
 hydrido and halo;

R_7 and R_{12} are independently selected from the group consisting of
 15 hydrido and halo.

34. The compound as recited in Claim 33 or a pharmaceutically
 acceptable salt thereof, wherein;

20 n is the integer 1;

R_1 is trifluoromethyl;

R_{16} is hydrido;

R_2 is hydrido;

R_3 is hydrido;

25 Y is methylene;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido and fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy, 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of cyclopentyl, 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of fluoro and hydrido;

R_7 and R_{12} are independently selected from the group consisting of hydrido and fluoro.

35. The compound as recited in Claim 34 or a pharmaceutically acceptable salt thereof, wherein;

n is the integer 1;

R_1 is trifluoromethyl;

R_{16} is hydrido;

R_2 is hydrido;

R_3 is hydrido;

Y is methylene;

Z is a covalent single bond;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group
5 consisting of hydrido and fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy,
4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy,
3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy,
3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy,
10 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy,
3-pentafluoroethylphenoxy, 3-tert-butylphenoxy,
3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl-2-yl)oxy,
3-trifluoromethoxybenzyl, 3-trifluoromethoxyphenoxy,
3-trifluoromethylbenzyl, and 3-trifluoromethylthiophenoxy;

15 R_{10} is selected from the group consisting of 1,1,2,2-tetrafluoroethoxy,
pentafluoroethyl, and trifluoromethyl;

R_6 and R_{11} are independently selected from the group consisting of
fluoro and hydrido;

R_7 and R_{12} are independently selected from the group consisting of
20 hydrido and fluoro.

36. A compound as recited in Claim 29 or a pharmaceutically acceptable salt
25 thereof wherein said compound is selected from the group consisting of:

- (2R)-3-[[3-(3-trifluoromethoxyphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-isopropylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[3-(3-cyclopropylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-(2-furyl)phenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2,3-dichlorophenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 10 (2R)-3-[[3-(4-fluorophenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-methylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-(2-fluoro-5-bromophenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-chloro-3-ethylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(1,1,2,2-tetrafluoroethoxy)phenoxy]phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[3-[3-(pentafluoroethyl)phenoxy]phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3,5-dimethylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(3-ethylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-*t*-butylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-methylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 30

- (2R)-3-[[3-(5,6,7,8-tetrahydro-2-naphthoxy)phenyl][[3-(1,1,2,2-tetrafluoro-ethoxy)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(phenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[3-[3-(*N,N*-dimethylamino)phenoxy]phenyl][[3-(1,1,2,2-tetrafluoro-ethoxy)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(1,1,2,2-tetrafluoroethoxy)phenyl]methyl][3-[[3-(trifluoromethoxy)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(1,1,2,2-tetrafluoroethoxy)phenyl]methyl][3-[[3-(trifluoro-10 methyl)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(1,1,2,2-tetrafluoroethoxy)phenyl]methyl][3-[[3,5-dimethylphenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(1,1,2,2-tetrafluoroethoxy)phenyl]methyl][3-[[3-(trifluoromethylthio)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[[3-(1,1,2,2-tetrafluoroethoxy)phenyl]methyl][3-[[3,5-difluorophenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 3-[[[3-(1,1,2,2-tetrafluoroethoxy)phenyl]methyl][3-[cyclohexylmethoxy]-phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 3-[[3-(2-difluoromethoxy-4-pyridyloxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[3-(2-trifluoromethyl-4-pyridyloxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-difluoromethoxyphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[[3-(3-trifluoromethylthio)phenoxy]phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-chloro-3-trifluoromethylphenoxy)phenyl][[3-(1,1,2,2-tetrafluoroethoxy)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-trifluoromethoxyphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 30

- (2R)-3-[[3-(3-isopropylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-cyclopropylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[3-(3-(2-furyl)phenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2,3-dichlorophenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-fluorophenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 10 (2R)-3-[[3-(4-methylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2-fluoro-5-bromophenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-(4-chloro-3-ethylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(1,1,2,2-tetrafluoroethoxy)phenoxy]phenyl][[3-(pentafluoroethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(pentafluoroethyl)phenoxy]phenyl][[3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[3-(3,5-dimethylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-ethylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-*t*-butylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(3-methylphenoxy)phenyl][[3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(5,6,7,8-tetrahydro-2-naphthoxy)phenyl][[3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 30

- (2R)-3-[[3-(phenoxy)phenyl][3-(pentafluoroethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(*N,N*-dimethylamino)phenoxy]phenyl][3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[3-(pentafluoroethyl)phenyl]methyl][3-[[3-(trifluoromethoxy)phenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(pentafluoroethyl)phenyl]methyl][3-[[3-(trifluoromethyl)-phenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(pentafluoroethyl)phenyl]methyl][3-[[3,5-dimethylphenyl]methoxy]-phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 10 (2R)-3-[[3-(pentafluoroethyl)phenyl]methyl][3-[[3-(trifluoromethylthio)phenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(pentafluoroethyl)phenyl]methyl][3-[[3,5-difluorophenyl]methoxy]-phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-(pentafluoroethyl)phenyl]methyl][3-[cyclohexylmethoxy]phenyl]-amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2-difluoromethoxy-4-pyridyloxy)phenyl][3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2-trifluoromethyl-4-pyridyloxy)phenyl][3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[3-(3-difluoromethoxyphenoxy)phenyl][3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-trifluoromethylthio)phenoxy]phenyl][3-(pentafluoroethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(4-chloro-3-trifluoromethylphenoxy)phenyl][3-(pentafluoroethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-trifluoromethoxyphenoxy)phenyl][3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 30 (2R)-3-[[3-(3-isopropylphenoxy)phenyl][3-(heptafluoropropyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;

- (2R)-3-[[3-(3-cyclopropylphenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-(2-furyl)phenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[3-(2,3-dichlorophenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-fluorophenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-methylphenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 10 (2R)-3-[[3-(2-fluoro-5-bromophenoxy)phenyl][[3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-chloro-3-ethylphenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-[3-(1,1,2,2-tetrafluoroethoxy)phenoxy]phenyl][[3-(heptafluoropropyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(pentafluoroethyl)phenoxy]phenyl][[3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3,5-dimethylphenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[3-(3-ethylphenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-*t*-butylphenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(3-methylphenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(5,6,7,8-tetrahydro-2-naphthoxy)phenyl][[3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(phenoxy)phenyl][[3-(heptafluoropropyl)phenyl]methyl]
- 30 amino]-1,1,1-trifluoro-2-propanol;

- (2R)-3-[[3-[3-(*N,N*-dimethylamino)phenoxy]phenyl][3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(heptafluoropropyl)phenyl]methyl][3-[[3-(trifluoromethoxy)phenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[[3-(heptafluoropropyl)phenyl]methyl][3-[[3-(trifluoromethyl)phenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(heptafluoropropyl)phenyl]methyl][3-[[3,5-dimethylphenyl]methoxy]-phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(heptafluoropropyl)phenyl]methyl][3-[[3-(trifluoromethylthio)phenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 10 (2R)-3-[[[3-(heptafluoropropyl)phenyl]methyl][3-[[3,5-difluorophenyl]methoxy]-phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(heptafluoropropyl)phenyl]methyl][3-[cyclohexylmethoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-(2-difluoromethoxy-4-pyridyloxy)phenyl][3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2-trifluoromethyl-4-pyridyloxy)phenyl][3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-difluoromethoxyphenoxy)phenyl][3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[[3-(3-trifluoromethylthio)phenoxy]phenyl][3-(heptafluoropropyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-chloro-3-trifluoromethylphenoxy)phenyl][3-(heptafluoropropyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(3-trifluoromethoxyphenoxy)phenyl][2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-isopropylphenoxy)phenyl][2-fluoro-5-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-cyclopropylphenoxy)phenyl][2-fluoro-5-
- 30 (trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;

- (2R)-3-[[3-(3-(2-furyl)phenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2,3-dichlorophenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[3-(4-fluorophenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-methylphenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2-fluoro-5-bromophenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 10 (2R)-3-[[3-(4-chloro-3-ethylphenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(1,1,2,2-tetrafluoroethoxy)phenoxy]phenyl][[2-fluoro-5-(trifluoro-methyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-[3-(pentafluoroethyl)phenoxy]phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3,5-dimethylphenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-ethylphenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[3-(3-*t*-butylphenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-methylphenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(5,6,7,8-tetrahydro-2-naphthoxy)phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(phenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(*N,N*-dimethylamino)phenoxy]phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 30

- (2R)-3-[[[2-fluoro-5-(trifluoromethyl)phenyl]methyl][3-[[3-(trifluoromethoxy)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[2-fluoro-5-(trifluoromethyl)phenyl]methyl][3-[[3-(trifluoromethyl)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[[2-fluoro-5-(trifluoromethyl)phenyl]methyl][3-[[3,5-dimethylphenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[2-fluoro-5-(trifluoromethyl)phenyl]methyl][3-[[3-(trifluoromethylthio)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[2-fluoro-5-(trifluoromethyl)phenyl]methyl][3-[[3,5-difluorophenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;
- 10 (2R)-3-[[[2-fluoro-5-(trifluoromethyl)phenyl]methyl][3-[cyclohexylmethoxy]-phenyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2-difluoromethoxy-4-pyridyloxy)phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-(2-trifluoromethyl-4-pyridyloxy)phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-difluoromethoxyphenoxy)phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[3-(3-trifluoromethylthio)phenoxy]phenyl][[2-fluoro-5-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 20 (2R)-3-[[3-(4-chloro-3-trifluoromethylphenoxy)phenyl][[2-fluoro-5-(trifluoro-methyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-trifluoromethoxyphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(3-isopropylphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-cyclopropylphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-(2-furyl)phenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 30

- (2R)-3-[[3-(2,3-dichlorophenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-fluorophenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- 5 (2R)-3-[[3-(4-methylphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(2-fluoro-5-bromophenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(4-chloro-3-ethylphenoxy)phenyl][[2-fluoro-4-
- 10 (trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(1,1,2,2-tetrafluoroethoxy)phenoxy]phenyl][[2-fluoro-4-(trifluoro-methyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(pentafluoroethyl)phenoxy]phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 15 (2R)-3-[[3-(3,5-dimethylphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]-methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-ethylphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-*t*-butylphenoxy)phenyl][[2-fluoro-4-
- 20 (trifluoromethyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(3-methylphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]methyl]-amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-(5,6,7,8-tetrahydro-2-naphthoxy)phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- 25 (2R)-3-[[3-(phenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[3-[3-(*N,N*-dimethylamino)phenoxy]phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;
- (2R)-3-[[[2-fluoro-4-(trifluoromethyl)phenyl]methyl][3-[[3-
- 30 (trifluoromethoxy)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;

(2R)-3-[[[2-fluoro-4-(trifluoromethyl)phenyl]methyl][3-[[3-(trifluoro-methyl)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;

(2R)-3-[[[2-fluoro-4-(trifluoromethyl)phenyl]methyl][3-[[3,5-dimethylphenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;

5 (2R)-3-[[[2-fluoro-4-(trifluoromethyl)phenyl]methyl][3-[[3-(trifluoromethylthio)-phenyl]methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;

(2R)-3-[[[2-fluoro-4-(trifluoromethyl)phenyl]methyl][3-[[3,5-difluorophenyl]-methoxy]phenyl]amino]-1,1,1-trifluoro-2-propanol;

10 (2R)-3-[[[2-fluoro-4-(trifluoromethyl)phenyl]methyl][3-[cyclohexylmethoxy]-phenyl]amino]-1,1,1-trifluoro-2-propanol;

(2R)-3-[[[3-(2-difluoromethoxy-4-pyridyloxy)phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;

(2R)-3-[[[3-(2-trifluoromethyl-4-pyridyloxy)phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;

15 (2R)-3-[[[3-(3-difluoromethoxyphenoxy)phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol;

(2R)-3-[[[3-(3-trifluoromethylthio)phenoxy]phenyl][[2-fluoro-4-(trifluoromethyl)-phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol; and

20 (2R)-3-[[[3-(4-chloro-3-trifluoromethylphenoxy)phenyl][[2-fluoro-4-(trifluoro-methyl)phenyl]methyl]amino]-1,1,1-trifluoro-2-propanol.

37. A pharmaceutical composition comprising a compound of one of claims 1 through 36 together with a pharmaceutically acceptable carrier.

25 38. A method of treating coronary artery disease or other CETP-mediated disorders in a subject by administering a therapeutically effective amount of a compound of one of claims 1 through 36.

30 39. A method of preventing coronary artery disease or other CETP-mediated disorders in a subject by administering a therapeutically effective amount of a compound of one of claims 1 through 36.

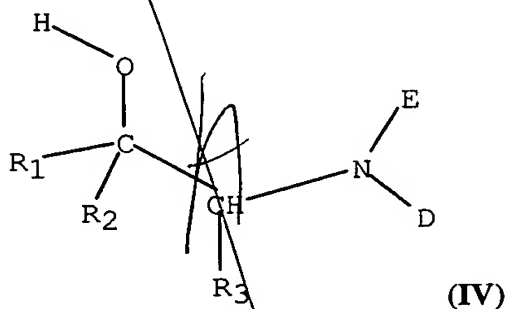
40. A method of preventing cerebral vascular accident (CVA) in a subject by administering a therapeutically effective amount of a compound of one of claims 1 through 36.

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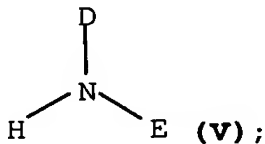
41. A method of preventing or treating dyslipidemia in a subject by administering a therapeutically effective amount of a compound of one of claims 1 through 36.

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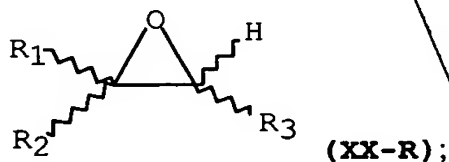
42. A process for the preparation of compounds as recited in any one of claims 1 or 2 having the Formula (IV):



and pharmaceutically acceptable salts thereof, comprising the reaction of an amine of Formula (V):



15 with an epoxide of Formula (XX-R):



wherein;

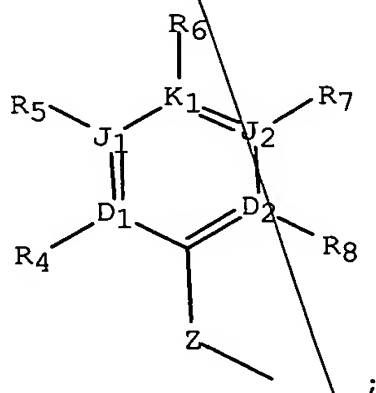
R_1 is selected from the group consisting of haloalkyl and haloalkoxymethyl;

R_2 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, haloalkoxy, haloalkoxyalkyl, perhaloaryl, perhaloaralkyl, perhaloaryloxyalkyl, and heteroaryl;

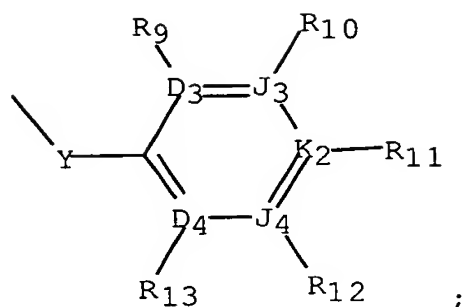
R_3 is selected from the group consisting of hydrido, aryl, alkyl, alkenyl, haloalkyl, and haloalkoxyalkyl;

D and E are independently selected from the group consisting of hydrido, A, and Q with the provisos that E and D are other than hydrido at the same time and A and Q are independently selected, when one of E and D is hydrido or when Y and Z are both single covalent bonds, from other than phenyl, 3-methylphenyl, 3-ethylphenyl, 2-methoxy-5-methylphenyl, 2-chlorophenyl, 3-chlorophenyl, and 3-bromophenyl;

A is the Formula:



Q is the Formula:



D_1, D_2, J_1, J_2 and K_1 are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that no more than one of D_1, D_2, J_1, J_2 and K_1 is a covalent bond, no more than one of D_1, D_2, J_1, J_2 and K_1 is O, no more than one of D_1, D_2, J_1, J_2 and K_1 is S, one of D_1, D_2, J_1, J_2 and K_1 must be a covalent bond when two of D_1, D_2, J_1, J_2 and K_1 are O and S, and no more than four of D_1, D_2, J_1, J_2 and K_1 are N;

D_3, D_4, J_3, J_4 and K_2 are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that no more than one is a covalent bond, no more than one of D_3, D_4, J_3, J_4 and K_2 is O, no more than one of D_3, D_4, J_3, J_4 and K_2 is S, no more than two of D_3, D_4, J_3, J_4 and K_2 are O and S, one of D_3, D_4, J_3, J_4 and K_2 must be a covalent bond when two of D_3, D_4, J_3, J_4 and K_2 are O and S, and no more than four of D_3, D_4, J_3, J_4 and K_2 are N;

Y is selected from the group consisting of a covalent single bond, $(CH_2)_q$ wherein q is an integer selected from 1 and 2, and $(CH_2)_j-O-(CH_2)_k$ wherein j and k are integers independently selected from 0 and 1;

Z is selected from the group consisting of covalent single bond,
 $(\text{CH}_2)_q$ wherein q is an integer selected from 1 and 2, and $(\text{CH}_2)_j\text{-O-}(\text{CH}_2)_k$
 wherein j and k are integers independently selected from 0 and 1;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group
 5 consisting of hydrido, halo, haloalkyl, and alkyl ;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the
 group consisting of hydrido, carboxy, heteroaralkylthio, heteroaralkoxy,
 cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy, heterocycloxy,
 aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocyclyl, perhaloaralkyl,
 10 aralkylsulfonyl, aralkylsulfonylalkyl, aralkylsulfinyl, aralkylsulfinylalkyl,
 halocycloalkyl, halocycloalkenyl, cycloalkylsulfinyl, cycloalkylsulfinylalkyl,
 cycloalkylsulfonyl, cycloalkylsulfonylalkyl, heteroaryl amino, N-
 heteroaryl amino-N-alkyl amino, heteroaryl aminoalkyl, haloalkylthio,
 alkanoyloxy, alkoxy, alkoxyalkyl, haloalkoxyalkyl, heteroaralkoxy,
 15 cycloalkoxy, cycloalkenyloxy, cycloalkoxyalkyl, cycloalkylalkoxy,
 cycloalkenyloxyalkyl, cycloalkylenedioxy, halocycloalkoxy,
 halocycloalkoxyalkyl, halocycloalkenyloxy, halocycloalkenyloxyalkyl,
 hydroxy, amino, thio, nitro, lower alkyl amino, alkylthio, alkylthioalkyl,
 aryl amino, aralkyl amino, arylthio, arylthioalkyl, heteroaralkoxyalkyl,
 20 alkylsulfinyl, alkylsulfinylalkyl, arylsulfinylalkyl, arylsulfonylalkyl,
 heteroaryl sulfinylalkyl, heteroaryl sulfonylalkyl, alkylsulfonyl,
 alkylsulfonylalkyl, haloalkylsulfinylalkyl, haloalkylsulfonylalkyl,
 alkylsulfonamido, alkylaminosulfonyl, amidosulfonyl, monoalkyl
 amidosulfonyl, dialkyl amidosulfonyl, monoaryl amidosulfonyl,
 25 arylsulfonamido, diarylamidosulfonyl, monoalkyl monoaryl amidosulfonyl,
 arylsulfinyl, arylsulfonyl, heteroarylthio, heteroaryl sulfinyl,
 heteroaryl sulfonyl, heterocyclylsulfonyl, heterocyclylthio, alkanoyl, alkenoyl,
 aroyl, heteroaroyl, aralkanoyl, heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl,
 alkynyl, alkenyloxy, alkenyloxyalkyl, alkylenedioxy, haloalkylenedioxy,

cycloalkyl, cycloalkylalkanoyl, cycloalkenyl, lower cycloalkylalkyl, lower cycloalkenylalkyl, halo, haloalkyl, haloalkenyl, haloalkoxy, hydroxyhaloalkyl, hydroxyaralkyl, hydroxyalkyl, hydroxyheteroaralkyl, haloalkoxyalkyl, aryl, heteroaralkynyl, aryloxy, aralkoxy, aryloxyalkyl, saturated heterocyclyl, partially saturated heterocyclyl, heteroaryl, heteroaryloxy, heteroaryloxyalkyl, heteroaralkyl, arylalkenyl, heteroarylalkenyl, carboxyalkyl, carboalkoxy, alkoxycarboxamido, alkylamidocarbonylamido, arylamidocarbonylamido, carboalkoxyalkyl, carboalkoxyalkenyl, carboaralkoxy, carboxamido, carboxamidoalkyl, cyano, carbohaloalkoxy, phosphono, phosphonoalkyl, diaralkoxyphosphono, and diaralkoxyphosphonoalkyl;

R_4 and R_5 , R_5 and R_6 , R_6 and R_7 , R_7 and R_8 , R_9 and R_{10} , R_{10} and R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} are independently selected to form spacer pairs wherein a spacer pair is taken together to form a linear moiety having from 3 through 6 contiguous atoms connecting the points of bonding of said spacer pair members to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 contiguous members, a partially saturated heterocyclyl ring having 5 through 8 contiguous members, a heteroaryl ring having 5 through 6 contiguous members, and an aryl with the provisos that no more than one of the group consisting of spacer pairs R_4 and R_5 , R_5 and R_6 , R_6 and R_7 , and R_7 and R_8 , is used at the same time and that no more than one of the group consisting of spacer pairs R_9 and R_{10} , R_{10} and R_{11} , R_{11} and R_{12} , and R_{12} and R_{13} is used at the same time.

43. A process according to Claim 42 wherein the reaction is carried out at a temperature of from 0 °C to 100 °C.

44. A process according to Claim 43 wherein the reaction is carried out at a temperature of from 15 °C to 65 °C.

45. A process according to Claim 42 wherein the process further comprises a solvent selected from the group consisting of tetrahydrofuran, dioxane,
5 methylene chloride, and acetonitrile.

46. A process according to Claim 42 wherein the process further comprises a transition metal salt catalyst selected from the group consisting of ytterbium, hafnium, scandium, neodymium, gadolinium, and zirconium salts.

47. A process according to Claim 46 wherein the transition metal salt is
10 selected from the group consisting of ytterbium triflate, hafnium triflate, scandium triflate, neodymium triflate, gadolinium triflate, and zirconium triflate.

48. A process according to Claim 42 wherein the oxirane XX-R is selected from the group consisting of 2-trifluoromethyloxirane,
15 2-pentafluoroethyloxirane, 2-(1,1,2,2-tetrafluoroethoxymethyl)oxirane, 2-(difluorochloromethyl)oxirane, and 2-(trifluoromethoxymethyl)oxirane.

49. A process according to Claim 42 in which the oxirane has the (R)-chiral configuration at the R₁ and R₂ substituted carbon.

20 50. A process according to Claim 42, wherein;

D₁, D₂, J₁, J₂ and K₁ are each carbon,

D₃, D₄, J₃, J₄ and K₂ are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that at least one
25 of D₃, D₄, J₃, J₄ and K₂ is selected from the group consisting of O, S, and N, wherein no more than one of D₃, D₄, J₃, J₄ and K₂ is a covalent bond, no

more than one of D_3 , D_4 , J_3 , J_4 and K_2 is O, no more than one of D_3 , D_4 , J_3 , J_4 and K_2 is S, one of D_3 , D_4 , J_3 , J_4 and K_2 must be a covalent bond when two of D_3 , D_4 , J_3 , J_4 and K_2 are O and S, and no more than four of D_3 , D_4 , J_3 , J_4 and K_2 are N.

5

51. A process according to Claim 42 wherein;

D_3 , D_4 , J_3 , J_4 and K_2 are each carbon;

D_1 , D_2 , J_1 , J_2 and K_1 are independently selected from the group consisting of C, N, O, S and covalent bond with the provisos that at least one of D_1 , D_2 , J_1 , J_2 and K_1 is selected from the group consisting of O, S, and N, wherein no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is a covalent bond, no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is O, no more than one of D_1 , D_2 , J_1 , J_2 and K_1 is S, one of D_1 , D_2 , J_1 , J_2 and K_1 must be a covalent bond when two of D_1 , D_2 , J_1 , J_2 and K_1 are O and S, and no more than four of D_1 , D_2 , J_1 , J_2 and K_1 are N.

15

52. A process according to Claim 42 wherein D_1 , D_2 , J_1 , J_2 , K_1 , D_3 , D_4 , J_3 , J_4 and K_2 are each carbon.

53. A process according to Claim 42 wherein;

D_1 , D_2 , J_1 , J_2 , K_1 , D_3 , D_4 , J_3 , J_4 and K_2 are each carbon

Y is selected from the group consisting of a covalent single bond and $(\text{CH}_2)_q$ wherein q is an integer selected from 1 and 2, and $(\text{CH}_2)_j\text{-O-}(\text{CH}_2)_k$ wherein j and k are integers independently selected from 0 and 1;

Z is a covalent single bond;

5

54. A process according to Claim 42 wherein;

$\text{D}_1, \text{D}_2, \text{J}_1, \text{J}_2, \text{K}_1, \text{D}_3, \text{D}_4, \text{J}_3, \text{J}_4$ and K_2 are each carbon

Y is selected from the group consisting of a covalent single bond and C1-C2 alkylene;

10

Z is a covalent single bond;

$\text{R}_4, \text{R}_8, \text{R}_9$, and R_{13} are independently selected from the group consisting of hydrido and halo;

15

$\text{R}_5, \text{R}_6, \text{R}_7, \text{R}_{10}, \text{R}_{11}$, and R_{12} are independently selected from the group consisting of hydrido, alkyl, halo, haloalkyl, haloalkoxy, aryl, alkylthio, arylamino, arylthio, aroyl, arylsulfonyl, aryloxy, aralkoxy, heteroaryloxy, alkoxy, aralkyl, cycloalkoxy, cycloalkylalkoxy, cycloalkylalkanoyl, heteroaryl, cycloalkyl, haloalkylthio, hydroxyhaloalkyl, heteroaralkoxy, heterocycloxy, aralkylaryl, heteroaryloxyalkyl, heteroarylthio, and heteroarylsulfonyl.

20

55. A process according to any of Claims 50 through 54 wherein the reaction is carried out at a temperature of from 0 °C to 100 °C.

56. A process according to Claim 55 wherein the reaction is carried out at a temperature of from 15 °C to 65 °C.

57. A process according to any of Claims 50 through 54 wherein the process further comprises a solvent selected from the group consisting of tetrahydrofuran, dioxane, methylene chloride, and acetonitrile.
58. A process according to any of Claims 50 through 54 wherein the process
5 further comprises a transition metal salt catalyst selected from the group consisting of ytterbium, hafnium, scandium, neodymium, gadolinium, and zirconium salts.
59. A process according to Claim 58 in which the transition metal salt is selected from the group consisting of ytterbium triflate, hafnium triflate,
10 scandium triflate, neodymium triflate, gadolinium triflate, and zirconium triflate.
60. A process according to any of Claims 50 through 54 wherein the oxirane XX-R is selected from the group consisting of 2-trifluoromethyloxirane, 2-pentafluoroethyloxirane, 2-(1,1,2,2-tetrafluoroethoxymethyl)oxirane,
15 2-(difluorochloromethyl)oxirane, and 2-(trifluoromethoxymethyl)oxirane.
61. A process according to Claim 60 in which the oxirane has the (R)-chiral configuration at the R_1 and R_2 substituted carbon.
62. A process according to any of Claims 50 through 54 wherein the process further comprises a:
- 20 (a) Temperature of from 0 °C to 100 °C;
(b) Non-protic solvent;
(c) Transition metal salt selected from the group consisting of ytterbium, hafnium, scandium, neodymium, gadolinium, and zirconium salts.
- 25 63. A process according to Claim 62 wherein the reaction is carried out at a temperature of from 15 °C to 65 °C.

64. A process according to Claim 62 wherein the solvent is selected from the group consisting of tetrahydrofuran, dioxane, methylene chloride, and acetonitrile.
- 5 65. A process according to Claim 62 wherein the transition metal salt is selected from the group consisting of ytterbium triflate, hafnium triflate, scandium triflate, neodymium triflate, gadolinium triflate, and zirconium triflate.
- 10 66. A process according to Claim 62 wherein the oxirane XX-R is selected from the group consisting of 2-trifluoromethyloxirane, 2-pentafluoroethyloxirane, 2-(1,1,2,2-tetrafluoroethoxymethyl)oxirane, 2-(difluorochloromethyl)oxirane, and 2-(trifluoromethoxymethyl)oxirane.
67. A process according to Claim 66 wherein the oxirane has the (R)-chiral configuration at the R_1 and R_2 substituted carbon.

add A³⁷